

**Structural and functional characterization of
natural alleles of potato (*Solanum tuberosum* L.)
invertases associated with tuber quality traits**

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**HIER MEIN GEHEIMNIS. ES IST GANZ EINFACH: MAN SIEHT NUR MIT
DEM HERZEN GUT. DAS WESENTLICHE IST FÜR DIE AUGEN
UNSICHTBAR...**

AUS DER KLEINE PRINZ VON ANTOINE DE SAINT-EXUPÉRY

ABSTRACT

The starch and sugar content of potato tubers are model traits for a candidate gene approach towards understanding the molecular basis of quantitative trait loci (QTL) because carbohydrate metabolism is one of the best studied plant processes at the functional level. Many genes of potato and other plant species involved in carbohydrate metabolism and transport have been cloned and functionally characterized. The starch and sugar content influences the nutritional quality of fresh potatoes and processed products, such as potato chips and French fries. Biochemical and genetic data suggest that genes encoding invertases (β -D-fructofuranosidase) participate in the control of starch and sugar content of the potato tubers. In this context, the focus was on invertases, which are ubiquitous enzymes hydrolyzing sucrose into the reducing sugars glucose and fructose. Under cold storage, reducing sugars accumulate in a genotype dependent manner in tubers (cold sweetening). High reducing sugar levels result in a reduction of potato chips quality e.g. darker chips colour, bitter taste, and a higher acrylamide concentration.

In this PhD work natural alleles of potato invertases, found previously to be significantly associated with tuber starch and sugar content (LI ET AL., 2005, 2008), were characterized at the molecular and biochemical level. Starting point was the PCR based cloning of cDNAs encoded by the five known potato invertase genes *Pain-1* on chromosome III, *invGE* and *invGF* on chromosome IX, and *pCD111* and *pCD141* on chromosome X, which resulted in 64 distinct alleles. Phenetic tree analysis based on amino acid similarity separated the alleles of cell wall and vacuolar invertases into two classes. This was also in line with the genomic organization of the different invertase loci. The genomic structures of the *Pain-1* locus and the gene pair *pCD111/pCD141* were unravelled by isolation and sequence analysis of corresponding BACs. The genes *pCD111* and *pCD141* consist each of six exons and five introns, and are arranged in a direct tandem repeat. The gene *Pain-1* comprises seven exons and six introns without a tandem duplication. Furthermore, structural alterations due to allelic invertase amino acid sequence variation were clarified by 3D-model analysis. Alleles associated with superior potato chips quality showed charge differences in the putative sucrose binding site possibly resulting in a lower conversion of sucrose in reducing sugars, which could influence potato chips quality. To test whether allele specific amino acid variations lead to altered enzyme activities, potato invertases were assayed in the yeast invertase mutant *SUC2*. Measurements of enzyme affinity to sucrose (K_m) and rate of sucrose conversion (v_{max}) did not show differences between vacuolar and cell wall-bound invertase alleles, which could explain the association with chips quality. Expression analysis revealed genotype specific transcription patterns of the alleles from different invertase isoforms.

In conclusion, at this point no definite statement on the functional impact of the characterized invertase alleles on potato chips quality is possible. The results of this study show that invertase regulation takes place at the transcript as well as at the protein level, is genotype specific and depends on allelic sequence variation. This work is a first example of a multi-pronged approach to dissect the natural variation of sugar and starch content in potato at molecular level.

Zusammenfassung

Stärke- und Zuckergehalt von Kartoffelknollen sind ausgezeichnete Beispiele, die molekularen Grundlagen von *quantitative trait loci* (QTLs) mittels Kandidatengenen zu identifizieren, da der Kohlenhydratstoffwechsel auf funktionaler Ebene einer der best untersuchten Prozesse in pflanzlichen Organismen ist. So wurden viele Gene aus Kartoffel und anderen Pflanzen, die mit dem Kohlenhydratstoffwechsel und -transport verbundenen sind, kloniert und hinsichtlich ihrer Funktion analysiert. Stärke- und Zuckergehalt beeinflussen unmittelbar die Qualität frischer Kartoffeln und weiter verarbeiteter Kartoffelprodukte wie zum Beispiel Kartoffelchips und Pommes Frites. Biochemische und genetische Daten deuten darauf hin, dass Invertase (β -D-fructofuranosidase) an der Kontrolle des Stärke- und Zuckergehaltes von Kartoffelknollen beteiligt ist. Deshalb wurde der Focus dieser Arbeit auf Invertasen gelegt, die universell verbreitet sind und Saccharose in die reduzierenden Zucker Glukose und Fruktose spalten. Eine durch Kaltlagerung bedingte Erhöhung an reduzierenden Zuckern in der Knolle ist abhängig von der Kartoffelsorte und äußert sich in einer Verschlechterung der Kartoffelchipsqualität, zum Beispiel dunklerer Farbe, bitterer Geschmack und einer höheren Konzentration von Acrylamid.

In dieser Doktorarbeit wurden natürlich vorkommende Allele von Kartoffelinvertasen, welche signifikant mit dem Stärke- und Zuckergehalt von Kartoffelknollen assoziiert sind (LI ET AL., 2005, 2008), hinsichtlich ihrer molekularen und biochemischen Eigenschaften charakterisiert. Die analysierten Gene waren *Pain-1* auf Chromosom III, *invGE* und *invGF* auf Chromosom IX, und *pCD111* und *pCD141* auf Chromosom X. Durch Klonieren von cDNA der fünf bekannten Invertasegene aus Kartoffel mittels PCR, konnten 64 Allele der unterschiedlichen Gene identifiziert werden. Weiterhin zeigte eine *phenetic tree*- Analyse, basierend auf Aminosäureähnlichkeiten der verschiedenen Invertaseisoformen, dass vakuoläre und apoplastische Invertasen in separate Klassen gegliedert werden können. Diese Einteilung wurde durch die unterschiedliche Genstruktur und -organisation der Isoformen, welche durch Isolation und Sequenzierung von entsprechenden BACs gewonnen wurde, bestätigt. So weisen die Gene *pCD111* und *pCD141* jeweils sechs Exons und fünf Introns auf und sind in einer direkten Tandem-Reihung angeordnet. Das Gen *Pain-1* enthält sieben Exons und sechs Introns, ohne ein Tandemduplikat aufzuweisen. Weiterhin wurden die durch allelische Aminosäuren ausgelösten strukturellen Veränderungen der Invertaseallele, mittels 3D-Analyse veranschaulicht. Allele, die nachweislich mit einer guten Kartoffelchipsqualität assoziiert sind, zeichneten sich durch Ladungsveränderungen in der potenziellen Saccharose-Bindedomäne des Enzyms aus. Diese Veränderungen führen möglicherweise zu einer verringerten Saccharosebindung und einer geringeren Konzentration an Glukose und Fruktose, was wiederum die Qualität von Kartoffelchips beeinflusst. Weiterführend wurde überprüft, ob allelspezifische

Aminosäurenvariationen zu einer veränderten Enzymaktivität führen können. Hierzu wurden die Enzymaffinität zu Saccharose (K_m) und die Rate der Saccharoseumwandlung (v_{max}) von Invertaseallelen aus Kartoffel mit Hilfe der Hefemutante *SUC2* analysiert. Die Messungen zeigten keine biochemischen Unterschiede zwischen apoplastischen und vakuolären Invertaseallelen, welche die Assoziation mit Kartoffelchipsqualität hätten erklären können.

Zusätzlich wurde mittels Expressionsanalyse veranschaulicht, dass allelische Transskriptionsmuster verschiedener Invertaseisoformen genotypspezifisch sind.

Zusammenfassend ist fest zu stellen, dass an diesem Punkt der Arbeit keine absolute Aussage über den funktionalen Einfluss der beschriebenen Invertaseallele auf Kartoffelchipsqualität möglich ist. Die Ergebnisse dieser Doktorarbeit zeigen, dass Invertaseregulation sowohl auf Transskript- als auch auf Proteinebene erfolgt, genotypspezifisch ist und von der Sequenzvariation der Allele abhängt. Diese Arbeit ist ein erstes Beispiel für einen ebenenübergreifenden Ansatz, die natürlich vorkommende Variation des Stärke- und Zuckergehalts in Kartoffelknollen auf molekularer Ebene zu entschlüsseln.

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Abbreviations

112 A1 NE	yeast expression vector
%	percent
°C	degree celsius
µg	microgram
µl	microliter
3'	three prime end of a DNA fragment
5'	five prime end of a DNA fragment
aa	amino acid
BAC	bacterial artificial chromosome
BNA	Böhm-Nordkartoffel Agrarproduktion GbR, 29574 Ebstorf
Btn	Biotin
bp	base pair(s)
ca.	circa
cDNA	complementary deoxyribonucleic acid
cM	centi Morgan
cm	centimeter
CQA	potato chips quality in autumn after harvest
CQS	potato chips quality in spring after 3-4 months of cold storage
DNA	deoxyribonucleic acid
DNAse	desoxyribonuclease
dNTP	deoxynucleosidetriphosphate
<i>E.coli</i>	<i>Escherichia coli</i>
e.g.	exempli gratia (Lat.) for example
EP	electrostatic potential
eQTLs	expression QTLs
EST	expressed sequence tag
<i>et al.</i>	<i>Et alii/et aliae</i> (Lat.) and others
EtBr	ethidium bromide
EtOH	ethanol
F1	first filial generation after cross
FAO	Food and agriculture organization of the United Nations
Fru	fructose
g	gram
Glc	glucose
GUS	β-glucoronidase
h	hour(s)
I	Inosin
InDels	insertions, deletions
INV	invertase
Inv-CW	cell wall-bound invertase
Inv-N	neutral invertase
Inv-V	vacuolar invertase

kb	kilo base pair(s)
kDa	kilo dalton
K _m	Michaelis constant
l	liter
LB medium	Luria Bertani medium
LD	linkage disequilibrium
M	molar
mg	milligram
min	minute(s)
mM	millimolar
mm	millimeter
MPIMP	Max-Planck-Institute for Molecular Plant Physiology/Golm
MPIZ	Max-Planck-Institute for Plant Breeding Research/Köln
mRNA	messenger ribonucleic acid
ng	nanogram
nm	nanometer
NOR	NORIKA GmbH, 18190 Groß Lüsewitz
OD	optical density
PAGE	polyacrylamide gel electrophoresis
PCR	polymerase chain reaction
PFGE	pulse field gel electrophoresis
pH	negative decimal log. of the H ⁺ concentration
pmol	picomolar
qRT-PCR	quantitative reverse transcription PCR
QTL	quantitative trait locus/loci
RFLP	restriction fragment length polymorphism
RNA	ribonucleic acid
rpm	rounds per minute
RT	room temperature
S.	Solanum
SAR	SAKA-RAGIS Pflanzenzucht GbR, Zuchtstation Windeby, 24340 Windeby/Eckernförde
SDS	sodium dodecil sulfate
sec	second(s)
SNP	single nucleotide polymorphisms
SSCP	single stranded conformation polymorphism
ssp.	subspecies
SUC	sucrose
TSC	tuber starch content
TSY	tuber starch yield
U	unit(s)
V	volt
v _{max}	maximal velocity
WT	wild type

Abbreviations for nucleic acids

A	Adenine
T	Thymine
C	Cytosine
G	Guanine

Abbreviations for amino acids

A	Alanine
C	Cysteine
D	Aspartate
E	Glutamate
F	Phenylalanine
G	Glycine
H	Histidine
I	Isoleucine
K	Lysine
L	Leucine
M	Methionine
N	Asparagine
P	Proline
Q	Glutamine
R	Arginine
S	Serine
T	Threonine
V	Valine
W	Tryptophan
Y	Tyrosine

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1 Introduction

1.1 The potato

The potato (*Solanum tuberosum*) originated in the highlands of South America, where it has been cultivated for more than 8,000 years. Spanish explorers introduced the plant into Europe in the 16th century as a botanical curiosity (BRÜCHER, 1975). By the 19th century it was cultivated throughout the continent, providing an abundant and inexpensive food source. Today the potato is the fourth most important food crop in the world following wheat, maize and rice (www.cipotato.org). More than one-third of the global potato output now comes from developing countries, comparing to just 11% in the early 1960s. According to the latest FAO data from 2007 (Food and agriculture organization of the United Nations; <http://faostat.fao.org>), potato production worldwide stands at 322 million tons and covers more than 19 million hectares. China is now the world's largest potato producer with 70 million tons yearly. Although potato production in Europe has fallen since the early 1960s, this decline has been more than offset by the growth in Asia, Africa, and Latin America, thereby explaining the global rise in global potato tonnage. Because of its importance, potato became an object for extensive genetic studies increasing its resistance to pathogens, as well as improving cooking and nutrition qualities like the starch/sugar content and potato chips quality.

The genus *Solanum* contains seven cultivated and 228 wild species. The potato genome occurs in a range of chromosome numbers from $2n=24$ (diploid) in for example. *S. stenotomum*, *S. phureja*, and *S. ajanhuiri*, $3n=36$ (triploid) in *S. caucha*, and *S. juzepczukii*, $4n=48$ in *S. tuberosum* ssp. *tuberosum* and *S. tuberosum* ssp. *andigena*, $5n=60$ (pentaploid) in *S. curtilobum* to $6n=72$ (hexaploid) in *S. demissum* (HAWKES, 1990).

The cultivated potato was named by Carl von Linné from a specimen grown in Europe and the scientific name *Solanum tuberosum* is presently used to include domesticated potatoes from South America, Europe and the USA and derivatives of them in the rest of the world (INGRAM & WILLIAMS, 1991).

The cultivated potato (*S. tuberosum* ssp. *tuberosum*) is far from being an ideal species for genetic analysis due to its tetraploidy with tetrasomic inheritance. In addition, the high heterozygosity, the owing to inbreeding depression after repeated selfing, and multiple allelism generating a multitude of genotypes that are difficult to distinguish by comparing expected with observed segregation ratios complicates molecular analysis. An allele at any given locus

can occur in four allelic states: simplex (present one time at the locus), duplex (present two times), triplex (present three times) and quadruplex (present four times, homozygous state). Therefore, at any given genetic locus, up to four different alleles are possible. Nevertheless, the use of molecular markers enables the selection of favourable genotypes at an early time point in the potato breeding process.

1.2 Crosstalk between starch and sugar metabolism and the phenomenon of cold sweetening

As described above, potato is economically highly important. Being a staple food, potato is not only grown as a vegetable for table use but also processed into French fries and potato chips and used for dried products and starch production. In developed countries, up to 66% of potato in everyday diet is consumed in processed form. Consumers demands for convenient food at home, fast food in restaurants and snacking gave rise to a wide variety of processed products. These include potato chips, French fries, and various other frozen products, dehydrated potato products, as well as chilled-peeled and canned potatoes. Potato chips are thinly sliced potatoes, which are fried at high temperatures in different types of vegetable oil. The biggest problem of potato chips quality is the consumer's preference on fry colour, which is 'light-golden'. Fry colour is strongly dependent upon the reducing sugar concentration in tubers. But also other factors like variety characteristics, maturity at harvest, and ethylene in the storage atmosphere contribute to potato chips quality. A high reducing sugar content in tubers causes major problems for food processing because high frying temperatures lead to the non-enzymatic Maillard reaction between free aldehyde groups of the reducing sugars glucose and fructose and free α -amino groups of amino acids and proteins (SHALLENBERGER ET AL., 1959). With increasing concentration of reducing sugars in the raw tuber, chips and French fries colour changes from light yellow to dark brown and the fried products get a bitter taste (ROE ET AL., 1990). This process also attracts further attention because of the discovery that acrylamide, a potential neurotoxin and group 2A potential carcinogen, is formed from asparagine and reducing sugars via an N-glycoside intermediate in a side step of the Maillard reaction (MOTTRAM ET AL., 2002; CHUDA ET AL., 2003; OHARA-TAKADA ET AL., 2005; MATSUURA-ENDO ET AL., 2006).

The accumulation of reducing sugars in plants at low temperatures is a widespread and well-established phenomenon, often referred to as cold-induced sweetening or cold sweetening (MÜLLER-THURGAU, 1882). In mature potato tubers, which are stored in the cold to prevent tuber sprouting and loss of moisture, a shift in the balance between starch degradation and

glycolysis occurs, which leads to sucrose accumulation. Sucrose is then enzymatically converted into the reducing sugars glucose and fructose, which interfere with potato chips quality.

The sugar and starch metabolism is one of the best studied processes in plants. In higher plants, starch and sucrose are the primary products of photosynthetic carbon fixation. Starch is synthesized as an intermediate deposit in the chloroplasts, and stored in non-photosynthetic organs in amyloplasts. Sucrose is synthesized in the cytosol, transiently stored in the vacuole and exported via the phloem to sink tissues. For long-term storage sucrose can be converted into polymers such as starch, triacyl glycerides, polypeptides, or secondary compounds, enabling plants to cope with environmental biotic or abiotic challenges (STURM ET AL., 1999). In most plants, sucrose is the major form how carbon is transported from source to sink organs because of the non-reducing nature of the disaccharide, in which glucose and fructose are linked ($\alpha 1 \rightarrow \beta 2$); (ARAI ET AL., 1991). Sucrose and the cleavage products glucose and fructose are the central molecules for carbohydrate translocation, metabolism, and sensing in higher plants (ROITSCH & GONZÁLEZ, 2004).

Starch may be degraded either hydrolytically or phosphorolytically (MALONE ET AL., 2006). The products are exported from the amyloplast either as hexose phosphates via the glucose phosphate-phosphate translocator or as free sugars via the glucose and/or maltose transporters (SMITH ET AL., 2005). Once in the cytosol, these metabolites are converted to sucrose via sucrose phosphate synthase (KRAUSE ET AL., 1998). Subsequently, a proportion of the sucrose may be hydrolysed to glucose and fructose by acid invertase (GREINER ET AL., 1999; WINTER & HUBER, 2000).

Several mechanisms have been proposed to explain the sugar accumulation during cold storage in plants (Figure 1.1). An early suggestion is that sweetening occurs because the entry of hexose phosphates into glycolysis is restricted at low temperatures. Therefore, the products of starch breakdown are directed into the pathway of sucrose synthesis (AP REES ET AL., 1981). Other studies implied that cold-induced sweetening requires long-term changes. ISHERWOOD ET AL. (1976) demonstrated that sugar accumulation is delayed several days following transfer to low temperatures. This delay may be associated with changes in gene expression involved in cold acclimation (VAN BUSKIRK & THOMASHOW, 2006).

A precise explanation how low temperatures may stimulate sugar accumulation is still unclear, but several possibilities have been suggested. One is that sugar accumulation requires the induction of starch degrading enzymes. It has been reported that enzyme activities of

endo- and exo-amylase (COCHRANE ET AL., 1991) and starch phosphorylase (CLAASSEN ET AL., 1993) were increased during cold storage leading to increased formation of hexose phosphates. However, this cold induction of starch degrading enzymes was not observed consistently because most studies failed to discriminate between different isoenzymes that might contribute to the overall amylolytic and phosphorylytic activity (MALONE ET AL., 2006). A second possibility is that sugar accumulation requires an increased activity of enzymes involved in sucrose synthesis. DEITING ET AL. (1998) showed a temperature dependence of a novel form of sucrose phosphate synthase accompanied by sugar accumulation. Another possibility suggests that the formation of glucose and fructose might be required to promote sugar accumulation by removing sucrose. DUPLESSIS ET AL. (1996) reviewed the potential of the so-called ‘alternative pathway’ (cyanine resistant respiration) in the starch to sucrose conversion. Low temperatures activate the alternative pathway, which leads to decreased ATP levels and simultaneous increase in sucrose concentration. This sucrose becomes the substrate for vacuolar acid invertase resulting in the accumulation of reducing sugars. However, the physiological sense of soluble sugar accumulation during cold temperatures is presumably frost protection for plant tissues (PRESSEY, 1966, 1969).

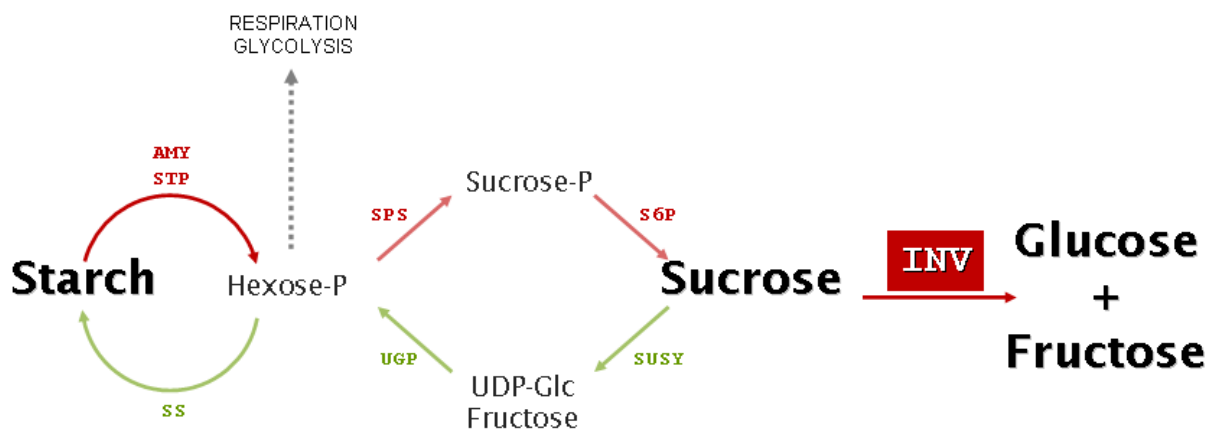


Figure 1.1: Simplified schematic overview of starch and sugar metabolism. Starch is degraded by endo- and exo-amylase (AMY) as well as by starch phosphorylase (STP). Products of these reactions are hexose phosphates (hexose-P), which can be shunted into glycolysis and respiration, or converted in starch by the action of starch synthase (SS). Another possibility is the synthesis of sucrose phosphate (sucrose-P) by sucrose phosphate synthase (SPS). Sucrose-P is then converted by sucrose 6-phosphatase (S6P) to sucrose. Sucrose can be degraded by sucrose synthase (SuSy) leading to the production of UDP-glucose and fructose. UDP-glucose is the substrate of UDP-glucose phosphatase leading to hexose-P. Another possibility of sucrose breakdown is the hydrolysis by invertases leading to the reducing sugars glucose and fructose, which mainly account for potato chips quality.

Cleavage of sucrose is mediated either by sucrose synthase (SuSy, EC 2.4.1.13) or by invertases (EC 3.2.1.26, β -fructosidase, β -fructofuranosidase). The reversible cleavage by sucrose synthase into UDP-glucose and fructose conserves the $\alpha 1 \rightarrow \beta 2$ glycosidic bond energy of sucrose. By contrast, invertases catalyze the irreversible hydrolysis yielding glucose

and fructose. The impact of SuSy and invertases in carbon partitioning are not clearly defined. Depending on the plant organ, tissue, and species, SuSy and invertases fulfil divergent functions and importance in the breakdown of sucrose. It is generally accepted that SuSy acts predominately in growing sink organs, whilst levels in photosynthetic source tissues are low (AP REES, 1984). In the developing potato tuber, the vast majority of sucrose degradation is catalyzed by SuSy, while invertases play only a significant role at the beginning of tuberization (APPELDOORN ET AL., 1997). The crucial role of SuSy in sucrose breakdown in tubers is its importance of sucrose mobilization for the synthesis of storage compounds as well as its predominance in determining the potato tuber sink strength (HEIM ET AL., 1993; SEBKOVÁ ET AL., 1993; ZRENNER ET AL., 1995).

Although studies suggest the major role of SuSy in sink strength regulation, also the participation of invertases in this process was demonstrated suggesting especially cell wall invertase isoforms as main determinants of sink strength. ROITSCH (1999) and ROITSCH ET AL. (2000) showed that cell wall invertase regulates sink strength especially during the initial stages of sink development. The essential role of cell wall-bound invertase isoforms in regulating phloem unloading and sink strength has been analysed in transgenic carrot plants (STURM & TANG 1999; TANG ET AL. 1999). Using antisense suppression of cell wall-bound invertases under the control of the 35S-CaMV promoter dominantly active in carrot tap roots resulted in lowered carbohydrate content in roots and increased leaf-to-root ratio. STITT & SONNEWALD (1995) demonstrated the induction of sink metabolism in source leaves of transgenic plants by over-expression of a yeast invertase. Another study showed that cell wall-bound invertase isoforms of maize play an important role in sucrose partitioning of the developing maize endosperm (MILLER & CHOUREY, 1992). In the developing seed coat of faba bean seeds, a correlation of the cell wall-bound invertases activity and high levels of hexoses in the cotyledons and in the apoplastic space has been determined (WEBER ET AL., 1995). Additionally, invertases serve as regulators of sucrose movement and utilization beyond developmental processes. As shown elsewhere they play an important role in pathogen defence responses (KOCAL ET AL., 2009) and cold protection (PRESSEY, 1966, 1969). In mature tubers invertase influences the accumulation of the reducing sugars glucose and fructose in response to cold storage. Several studies demonstrated the influence of low temperature on increasing invertase activity (ROREM & SCHWIMMER, 1963; PRESSEY & SHAW, 1966) and invertase transcript up-regulation (ZRENNER ET AL., 1996; ZHOU ET AL., 2004; BAGNARESI ET AL., 2008).

1.3 Invertases

As described above, the accumulation of reducing sugars in mature tubers is due to invertase activity rather than to SuSy induced sucrose cleavage.

Invertases are ubiquitous enzymes in plants and occur in several isoforms. Based on their solubility, subcellular localization, pH optimum, and isoelectric point, three different types of invertase isoenzymes can be distinguished: vacuolar, cell wall-bound, and neutral invertases (Figure 1.2).

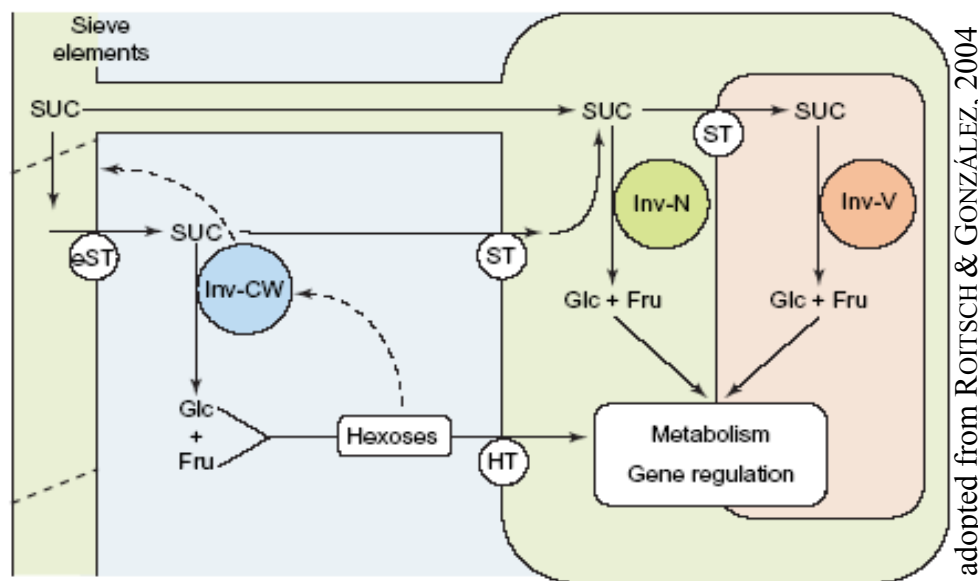


Figure 1.2: Subcellular location of invertase isoenzymes and phloem unloading pathways. Sucrose can be hydrolysed by different invertase isoforms depending on the mechanism of phloem unloading in the sink tissue. In symplastically isolated tissues, sucrose is unloaded from the sieve elements of the phloem into the apoplast by an assumed efflux sucrose transporter (eST). Once arrived in the apoplast, sucrose can be cleaved by a cell wall-bound invertase (Inv-CW). The resulting hexoses will be transported into the sink cell by a hexose transporter (HT). Alternatively, sucrose can be transported into the sink cells by a sucrose transporter (ST). Sucrose can be unloaded in the sink cells through the apoplast or by plasmodesmata. In the cytosol sucrose can be cleaved by neutral invertase (Inv-N) or sucrose synthase (SuSy). Sucrose is also stored in the vacuole for sucrose utilization and remobilization for metabolic processes. In the vacuole sucrose can be hydrolysed by vacuolar invertase (Inv-V). The hexoses glucose and fructose, which resulted from sucrose cleavage, are not only substrates for heterotrophic growth but also function as regulators of gene expression. Invertases also play a role in sink strength regulation maintaining and stimulating the enhancement of the flow of assimilates from source to sink. Abbreviations: SUC=sucrose, Glc=glucose, Fru=fructose.

Vacuolar and cell wall-bound invertases have similar enzymatic and biochemical properties and share two conserved amino acid motifs. Both types of invertases are β -fructofuranosidases, which act in an acidic pH optimum and are able to convert other fructofuranosidases such as stachyose and raffinose. The conversion of alternative substrates is less efficient than sucrose cleavage. Another interesting fact is that vacuolar and cell wall-bound invertases are glycoproteins (STOMMEL & SIMON, 1990). Glycosylation of vacuolar and extracellular glycoproteins is fairly different. Most of the vacuolar glycoproteins described so

far were found to be N-glycosylated with modified N-glycans containing fucose and/or xylose residues, but without terminal glucosamine residues. In contrast, extracellular glycoproteins were found to be N-glycosylated mostly by complex-type N-glycans including large structures with terminal fucose and galactose residues (RAYON ET AL., 1998). Both types of glycoproteins are post-Golgi modified, leading to protein maturation in the vacuole or in the extracellular compartment. It was shown that N-glycosylation in plants plays an important role in the induction of correct folding, the biological activity of the protein, and prevention of proteolytic degradation (reviewed in RAYON ET AL., 1998). FAYE & CHRISPEELS (1989) demonstrated the proteolytic degradation of unglycosylated cell wall-bound invertase of carrot in the secretory pathway or immediately after its arrival in the cell wall. Furthermore, N-linked oligosaccharides may contain targeting information or may be involved in protein recognition (RAYON ET AL., 1998). Glycosylation of acid invertases is required for their transport across either the plasma membrane or the tonoplast, according their localization in the apoplast or the vacuole.

Invertase isoenzymes are encoded by small gene families. The genomic organization and exon/intron structure seems to be conserved between monocotyledons and dicotyledons (reviewed in TYMOWSKA-LALANNE & KREIS, 1998 and ROITSCH & GONZÁLES, 2004). Plant invertases isolated to date have fairly similar structures and consist of six to eight exons. A special feature is the presence of a mini exon of only nine nucleotides that was found in all invertase genes except *InvDC1* of carrot (RAMLOCH-LORENZ ET AL., 1993; LORENZ ET AL., 1995; SIMPSON ET AL., 2000). The mini exon, one of the smallest known, encodes the three amino acids DPN of the highly conserved β -fructosidase motif N-DNP-G/A that spans three exons and is found in invertases from plants, bacteria, and yeasts (reviewed in ROITSCH & GONZÁLES, 2004). BOURNAY ET AL. (1996) observed that under cold stress the mini exon of the *pCD111* potato cell wall invertase was skipped in an alternative splicing event. The functional relevance of the splicing effect was not investigated.

Acid invertases share another conserved peptide domain, which belongs to the active site of the enzyme. Within this domain WECxDF x stands for yaline in vacuolar invertases while it is proline in cell wall-bound invertases (periplasmatic), (TYMOWSKA-LALANNE & KREIS, 1998; STURM, 1999). This conserved single amino acid difference determines the distinctly more acidic pH optimum of cell wall-bound invertases and their substrate specificity (GOETZ & ROITSCH, 1999).

Vacuolar and cell wall-bound invertases are synthesized as preproteins with a leading sequence that is cleaved off during transport and protein maturation. The leader sequence

separates in two fragments, a signal peptide and an N-terminal extension. The signal peptide is required for entry into the endoplasmic reticulum (ER) and consists of a basic region, a hydrophobic core of approximately 20 amino acids, and a polar region. The N-terminal extension is characteristic for all known vacuolar invertases and might be involved in the regulation of enzymatic activity, folding and stability of the polyprotein, or in vacuolar targeting (reviewed in TYMOWSKA-LALANNE & KREIS, 1998). Additionally, vacuolar invertases contain a C-terminal extension, not present in cell wall-bound invertases. This extension might be also involved in vacuolar targeting of the enzyme (BEDNAREK & RAIKHEL, 1992). Mature invertases show a molecular mass in the range of 60kDa.

Vacuolar invertases, also referred to as soluble acid invertases, are characterized by an acidic pH optimum of 5.0-5.5 and are localized in the vacuole (BOLLER & KENDE, 1979). They are crucial in determining the level of sucrose stored in the vacuole and in remobilization of sucrose for metabolic processes. Another well studied function is the regulation of the sugar balance in fruit tissues of tomato and mature potato tubers (OHYAMA et al., 1995; SCHOLES ET AL., 1996; GREINER ET AL., 1998). In potato one locus on chromosome III is known to encode vacuolar invertase. This locus is named *Pain-I*. To date, no information about genomic sequence, exon/intron structure and gene organization is available. Searching for orthologous genes in the syntenic tomato genome resulted in one EST on tomato chromosome III (www.sgn.cornell.edu). The activity of vacuolar invertases is correlated with the hexose/sucrose ratio in cold stored tubers (PRESSEY & SHAW, 1966), which leads to an increased accumulation of glucose and fructose in mature tubers. However, RICHARDSON ET AL. (1990) showed that there is no correlation between invertase activity and the total amount of accumulated sugars. Similarly, while antisense inhibition of invertase results in a decrease in the amount of hexoses and an increase in the amount of sucrose it does not affect the total amount of sugars accumulated (ZRENNER ET AL., 1996). The reducing sugars, increased during tuber cold storage due to cold protection, interfere with the trait potato chips quality that is analyzed in this study. The gene *Pain-I* is known to be differentially expressed during tuber cold storage (ZRENNER ET AL., 1996; ZHOU ET AL., 2004; BAGNARESI ET AL., 2008), showing transcript accumulation under the latter conditions.

Cell wall-bound invertases, also referred to as insoluble acid invertases, extracellular and periplasmatic invertases, are characterized by a low pH optimum of 3.5-5.0. They are localized in the apoplast and ionically bound to the cell wall. Cell wall-bound invertases cleave sucrose on its way from sieve elements of the phloem to the apoplast. The cleavage products glucose and fructose are then transported into the sink cells by hexose transporters.

In potato two different loci encoding cell wall-bound invertases are known. The first locus is *Inv_{ap}-b* on chromosome IX, which consists of two invertase genes *invGE* and *invGF* linked in a direct tandem repeat and separated by approximately 2.3kb from each other (MADDISON ET AL., 1999). The locus has a size of approximately 8.6kb. The genes *invGE* and *invGF* exhibit a similar exon/intron structure composed of six exons and five introns. The second locus of cell wall-bound invertases is *Inv_{ap}-a* on chromosome X, which consist of two genes *pCD111* and *pCD141*. Comparative analysis of the potato genetic map revealed that the gene pair *pCD111/pCD141* on chromosome X arose by partial chromosome duplication of the gene pair *invGE/invGF* of chromosome IX, and, therefore, might also be organized in a direct tandem repeat (GEBHARDT ET AL., 2003; FRIDMAN ET AL., 2003).

The third type of invertases are neutral invertases, also known as alkaline or cytoplasmatic invertases because of their pH optimum between 6.8 and 8.0, and their localization in the cytoplasm. Limited information about the physiological function is available due to the labile nature of this invertase isoform and low enzyme activity. In contrast to acidic invertases, neutral invertases are not glycosylated and preferentially or solely hydrolyse sucrose. Therefore, they are no fructofuranosidases, which also convert substrates such as stachyose and raffinose like described for vacuolar and cell wall-bound invertase isoenzymes. Neutral invertases are strongly inhibited by their cleavage products but not by heavy metals. The latter observation suggests marked differences in the catalytic site compared to acidic invertases. Neutral invertases were only found in cyanobacteria and plants suggesting an origin from orthologous prokaryotic genes after endosymbiosis (VARGAS ET AL., 2003). Native polypeptides of neutral invertase are homotetramers composed of subunits with a molecular mass of 54 to 65kDa (ROSS ET AL., 1996), except the enzyme of carrot (LEE & STURM, 1996). In potato no locus for neutral invertase was detected so far and no sequence information is available.

1.4 The genetic map of potato

The potato map is one of the most highly marker-saturated maps among the crop plant species. Initially, two potato maps were constructed concurrently using RFLP markers on different genetic backgrounds (BONIERBALE ET AL., 1988; GEBHARDT ET AL., 1989). These maps were then compared with each other and also aligned to the tomato RFLP map (GEBHARDT ET AL., 1991; TANKSLEY ET AL., 1992). Comparative mapping revealed that the genomes of potato and tomato are co-linear except for paracentric inversions of five chromosome arms (BONIERBALE ET AL., 1988; TANKSLEY ET AL., 1992). Comparison of the potato genetic map with the physical map of the sequenced *Arabidopsis* genome displayed syntenic relationships between circa 40% of the potato genetic map and circa 50% of the physical map of this very distantly related plant species (GEBHARDT ET AL., 2003). With the development of new molecular markers, the potato map was enriched, and at the moment it is based on more than 350 markers, which cover approximately 90% of the potato genome (GEBHARDT & VALKONEN, 2001). Molecular markers originate from natural DNA variation present in a population of individuals of the same species. The molecular basis of the variations are point mutations (SNP, single nucleotide polymorphisms), and insertions, deletions (InDels), or inversions of DNA fragments in one allele versus another.

The existence of highly marker saturated potato maps allows localizing many genes on the twelve potato chromosomes, and markers linked to these genes can be used to perform positive marker-assisted selection. Potato function maps have been constructed for pathogen resistance (GEBHARDT & VALKONEN, 2001), and for tuber traits e.g. tuber starch content (FREYRE & DOUCHES, 1994; SCHÄFER-PREGL ET AL., 1998), chips colour (FREYRE & DOUCHES, 1994), and cold sweetening (MENÉNDEZ ET AL., 2002), and many more traits (reviewed in GEBHARDT ET AL., 2004). Integration of positional information on genetic factors controlling agronomic characteristics results in potato function maps as a basis for innovative approaches improving breeding. Understanding the molecular basis of complex traits and the underlying genes and proteins will enable combination of favourable alleles in improvement programs, leading to ‘precision breeding’.

❖ Quantitative trait loci (QTL)

The starch and sugar content of potato tubers are quantitative traits. These physiological characteristics show continuous or quantitative phenotypic variation because phenotypic expression is controlled by more than one gene. Additionally, environment has a large influence. The precise number of the genes involved in a quantitative trait is usually not

known. The loci where such genes are located in the genome are referred to ‘quantitative trait loci’ (QTL), (GELDERMANN, 1975). The traits can be genetically dissected using linkage maps that are based on molecular markers (reviewed by TANKSLEY, 1993).

Structures and functions of proteins encoded by a QTL are unknown. It is assumed that DNA polymorphisms must exist in the gene(s) responsible for the observed QTL effect. These DNA polymorphisms are the molecular basis for phenotypic selection of superior genotypes by breeding. In contrast to many mutations with drastic effects on the phenotype, the molecular and functional variability present in genes controlling QTL operates under field conditions and, therefore, should not have severe effects on fitness.

QTL maps are the first step toward the identification of the responsible genes, either by positional cloning or by candidate gene approach.

QTLs for tuber starch and sugar content or chips colour have been mapped in potato (DOUCHES & FREYRE, 1994; SCHÄFER-PREGL ET AL., 1998; MENÉNDEZ ET AL., 2002). A number of candidate genes have been identified regarding the fact of co-localisation with QTLs or their position on molecular maps, as well as being functional in the biosynthesis, degradation, or transport of starch and sugars in potato and other plants (CHEN ET AL., 2001; MENÉNDEZ ET AL., 2002). Among others, invertase genes were identified as positional candidates for cold-sweetening QTLs.

❖ Association genetics based on candidate genes

Association genetics based on candidate genes is one approach aiming towards the improvement of crop breeding programs (‘precision breeding’). The candidate gene approach is based on the knowledge of a gene’s function controlling the trait of interest. Additionally, co-localization of a functional candidate gene with a QTL for the trait of interest leads to a positional candidate (PFLIEGER ET AL., 2001).

Association analysis was performed for potato vacuolar invertase and cell wall-bound invertase isoforms due to the fact of being functional candidates in the tuber sugar metabolism as well as their co-localization with QTLs for tuber sugar content (CHEN ET AL., 2001; MENÉNDEZ ET AL., 2002).

The locus *Pain-1* on chromosome III codes for a vacuolar invertase isoform. Association analysis of allelic *Pain-1* fragments revealed strong associations of the SSCP fragment *Pain1-9a* (8c, 5c) with potato chips quality after cold storage, explaining 10.4% of the phenotypic variance (LI ET AL., 2008). This marker allele also showed association with chips quality without cold storage, tuber starch content and tuber starch yield (Table 1.1). Furthermore, a

negative association of the allelic *Pain-1* fragment *Pain1-5b* with potato chips quality after cold storage was detected, which explains 2.5% of the phenotypic variance. This SSCP fragment is also negatively associated with tuber starch content (Table 1.1).

In addition, association analysis was carried out for the genes *invGE*, *invGF* and *pCD141*, which encode cell wall-bound invertase isoforms. LI ET AL. (2005, 2008) showed that *invGE* and *invGF* allelic fragments, which were in linkage disequilibrium, were associated with better potato chips quality. The SSCP fragments *invGE-6f* and *invGF-4d* explain 2.8% of phenotypic variance of potato chips quality after tuber cold storage (Table 1.1). Additionally, these alleles displayed association with better potato chips quality without cold storage. The genes *invGE* and *invGF* co-localize with a QTL for starch and sugar content of potato tubers *Sug9a* (CHEN ET AL., 2001; MENÉNDEZ ET AL., 2002).

Analysis of allelic fragments of the gene *pCD141* revealed a negative association of one SSCP fragment *pCD141-3c* with potato chips quality without cold storage, after storage in the cold and tuber starch yield (LI ET al., 2008; Table 1.1).

Table 1.1: Invertase-trait association of tetraploid potato genotypes (Li et al., 2008).

Invertase genes	Chr. no.	Marker allele	CQA $q (R^2)$	CQS $q (R^2)$	TSC $q (R^2)$	TSY $q (R^2)$
<i>Pain-1</i>	III	<i>Pain1-5b</i> <i>Pain1-9a</i> (8c, 5c)	ns 0.001 (4.4) ↑	0.048 (2.5) ↓ 0.000 (10.4) ↑	0.002 (5.3) ↓ 0.000 (12.0) ↑	ns 0.003(5.3) ↑
<i>invGE</i> / <i>invGF</i>	IX	<i>invGE-6f</i> (<i>invGF-4d</i>)	0.034 (2.2) ↑	0.040 (2.8) ↑	ns	ns
<i>pCD111</i> / <i>pCD141</i>	X	<i>pCD141-3c</i>	0.008 (3.1) ↓	0.028 (3.1) ↓	0.007 (4.2) ↓	ns

Associations $q > 0.05$ are significant, ns=not significant ($q < 0.05$). The amount of variance (in %) explained by the marker allele is given by the R^2 statistic. Marker fragments shown in parenthesis are in nearly absolute LD, having identical or highly similar distribution in the population and show similar associations. ↑↓: direction of effect. ↑ the marker allele has a positive effect on the trait. ↓ the marker allele has a negative effect on the trait.

Association of one marker allele with more than one trait might be explained by the involvement of the same gene in multiple metabolic pathways interconnecting these complex traits. The pleiotropic effects of individual alleles always have the same direction, either positive (more starch, less reducing sugars, better potato chips quality) or negative (less starch, more reducing sugars, inferior chips quality).

1.5 Objectives of this thesis

Based on the previous studies of QTL detection for potato tuber starch and sugar content as well as association analysis of candidate genes of the starch and sugar metabolism, invertases were found to fulfil criteria being positional and functional candidates. Objective of this study was the identification and characterization of allelic variation of vacuolar and cell wall-bound invertase isoforms contributing to the important agronomic trait potato chips quality. The project aims can be summarized as follows (i) identification of alleles of all three known potato invertase loci *Pain-1*, *Inv_{ap}-b* and *Inv_{ap}-a*; (ii) analysis of putative effects of allele specific amino acid differences using 3D structural comparison; (iii) biochemical analysis of invertase alleles and (iv) allele specific differential expression analysis.

Figure 1.3 shows the aspects dealt with in each of the following chapters.

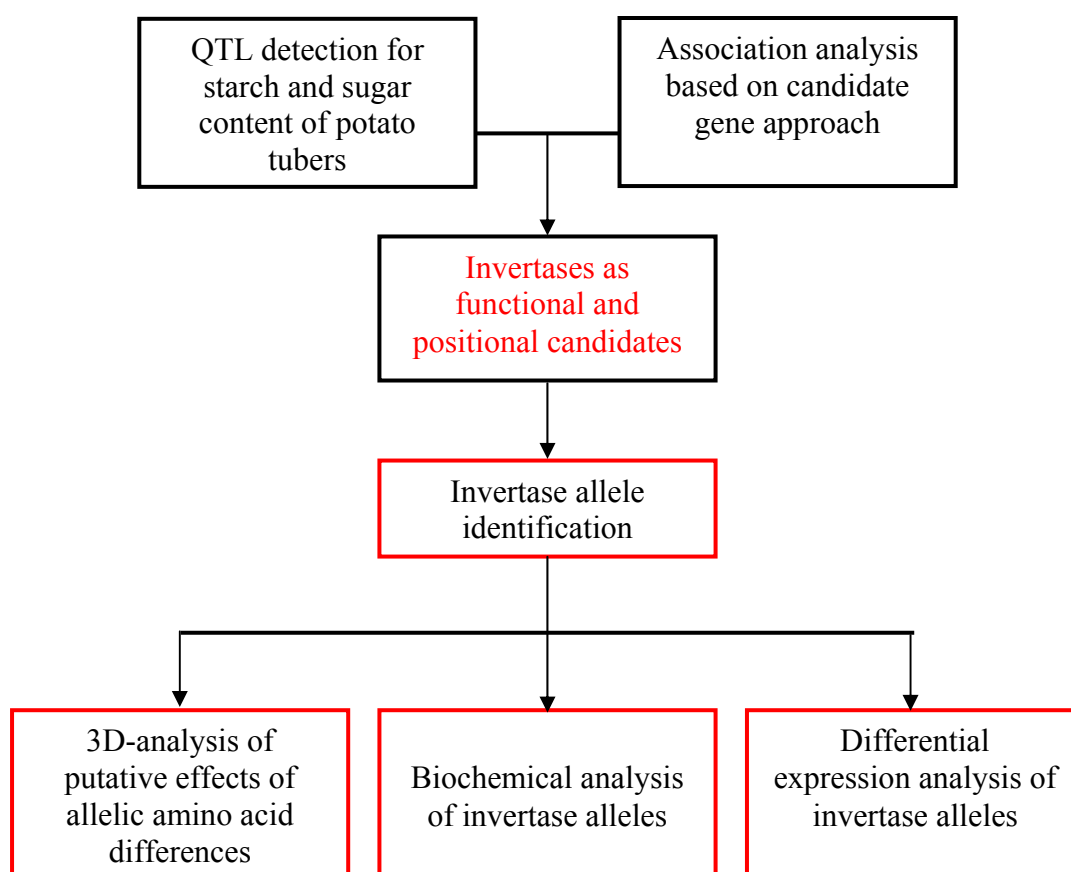


Figure 1.3: Strategy of the PhD work. The black boxes contain aspects, for which information was available. The red boxes contain aims of the study, which were set to elucidate the impact of different alleles of invertase isoforms on the analyzed trait potato chips quality.

The present study pursues to elucidate whether natural allelic variation of functional potato alleles within different genotypes can be identified and to what extent this natural variation

accounts for the observed phenotypic diversity of the selected genotypes regarding the trait chips quality. In consequence, the question arises whether allelic variation manifests at the functional level resulting in phenotypic diversity and whether these functional differences can be characterized, like observed in tomato for the fruit sugar content (FRIDMAN ET AL., 2004). This work will show whether natural variation of potato invertases is responsible for part of the phenotypic variation of tuber starch and sugar content, ultimately of potato chips quality.

2 Materials and Methods

2.1 Materials

2.1.1 Chemicals and Antibiotics

The chemicals and antibiotics (Table 2.1.1) were purchased from the following suppliers: Amersham Pharmacia Biotech (Braunschweig), Carl Roth GmbH (Karlsruhe), Difco Laboratories (Detroit, Michigan, USA), Invitrogen GmbH (Karlsruhe), Merck, Feinchemikalien und Laborbedarf (Darmstadt), Roche (Mannheim), and Sigma-Aldrich Chemie GmbH (Taufkirchen).

Table 2.1.1: Antibiotics.

Antibiotics	Solvent	Final conc. for selection on LB
Ampicillin (Amp)	H ₂ O	100µg/ml
Chloramphenicol (Cam)	Ethanol	12,5mg/l
Tetracyclin (Tet)	H ₂ O	12,5mg/l

2.1.2 Buffers and Culture Media

Stock solutions of the following buffers and culture media were prepared as described by SAMBROOK & RUSSELL (2001): DNA loading buffer, SDS, TAE, TBS, LB, YPD and SC medium. All media and solutions were made with highly purified Milli-Q-water (Millipore, Bedford, USA). Whenever required, the solutions were autoclaved for 20min at 121°C.

2.1.3 Restriction enzymes, nucleic acid modifying enzymes and kits

Restriction enzymes and corresponding buffers were used from New England BioLabs GmbH (Schwalbach/Taunus), MBI Fermentas GmbH (St. Leon-Rot) and Roche (Mannheim).

Nucleic modifying enzymes were used from:

DNA-freeTM Kit (Ambion, Cambridgeshire, UK)

Fast Start High Fidelity PCR System (Roche, Mannheim)

KOD Hot Start DNA Polymerase (Novagen, Darmstadt)

Ribonuclease Inhibitor (Roche, Mannheim)

RNase H (Roche, Mannheim)

SuperscriptTM II reverse transcriptase (Invitrogen, Karlsruhe)

Taq DNA Polymerase (PLUTHERO, 1993)

T4 DNA ligase (Invitrogen, Karlsruhe)

The following commercial reagents and kits were used:

Bio-Safe™ Coomassie G-250 stain (Bio-Rad, Hercules, USA)

ExoSAP-IT® (USB Corporation, Cleveland, USA)

First Strand cDNA Synthesis kit (Invitrogen, Karlsruhe)

High Pure PCR Purification kit (Roche, Mannheim)

LiCrosolv® (Merck KGaA, Darmstadt)

Miniprep® Kit/Midiprep® (Qiagen, Hilden)

P3504 Ponceau S (Sigma-Aldrich Chemie GmbH, Taufkirchen)

pGEM® - T and pGEM® - T Easy Vectors (Promega, Mannheim)

Power SYBR® Green PCR Master Mix (Applied Biosystems, Carlsbad, USA)

Protein Assay (Bio-Rad, Hercules, USA)

PSQ 96 SNP Reagent Kit (Biotage AB, Uppsala, Sweden)

PureLink™ Plant RNA Reagent (Invitrogen, Karlsruhe)

RNAwiz™ (Ambion, Cambridgeshire, UK)

Streptavidin-coated Super Paramagnetic beads (Amersham Biosciences, Sweden)

ToTally RNA™ (Ambion, Cambridgeshire, UK)

2.1.4 Oligonucleotides

Synthetic oligonucleotides were purchased from Invitrogen (Karlsruhe), Sigma-Aldrich Chemie GmbH (Taufkirchen), and Operon Biotechnologies GmbH (Köln). The primers used in this study are listed in the following Tables.

Table 2.1.2: Primers used for molecular cloning of PCR products.

Name	Primer sequence in 5'-3' orientation
ZrPain-F	ATGGCCACGCAGTACC
PainUni-R	GATGAATTACAAGTCTTGCAAGGG
PainNotI_F	CCCCGCGGCCGCATGGCCACGCAGTACC
PainNotI_R	ATATGCGGCCGCGATGAATTACAAGTCTGG
PainBamHI_R	CCCCGATCCGATGAATTACAAGTCTTGCAAGGG
PainGeno_For	GGGATTAACATGAGATCGTGTG
PainGeno_Rev	CCTGTACAGATGCCCATCCC
invGE-F-fulgth	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGGT
invGE-R-fulgth	TTAGTGCATCTTAGGTACATCCATGCTCCAAGC
ENotI_F	GATTGCGGCCGCATGGAATTATTTATGAAAAGC
ENotI_R	CCCCGCGGCCGCTTAGTGCATCTTAGGTACATCC
invGF-F-fulgth	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAG
invGF-R-fulgth	TCAATATTGTATCTTAGCTTTGCCATACTCCATGC
FNotI_F	CCCCGCGGCCGCATGGATTATTCATCTAATTC
FBamHI_R	CCCCGATCCTCAATATTGTATCTTAGCTTTGCCC
CD111fl_F	ATGGATTGTTTAAAAAAGTCTTCTC
CD111fl_R	TCAATAAGAAGAGTGACCAAATGACCAATTCA
CD141fl_F	ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTT
CD141fl_R	CTAGTGCAACTTTCATTAGCCATGCTCCAAGC

Table 2.1.3: Primers used for colony-PCR and for sequencing of constructs and PCR products.

Name	Primer sequence in 5'-3' orientation
T7	GTAATACGACTCACTATAGGGC
SP6P	TATTTAGGTGACACTATAG
Adh1-Prom	CTCACCATATCCGCAATGAC
Adh1-Term	CTTGAGTAACTCTTTCCTGTAGGTC
G112_1	GGTCAGGTTGCTTCTCAGGTATAG
G112_2	GAGAGTAGTGTGCGTGAATGAAGG
G112_3	CAGAACAGAAATGCAACGCG
G112_4	CCCAGTCACGACGTTGTAAAAC
G112_5	CCAACAAAGAATCTATACTTC
G112_6	CGACGCTCAAGTCAGAGGTGG
G112_7	GCTCGTTACAGTCCGGTGCG
G112_8	CATTGCGGATATGGTGAGACAAC
G112_9	GCGCATCACCAACATTTTCTG
G112_10	CTGACAGTTACCAATGCTTAATC
PGo2	GCCTCCCATTACACATTCTC
PGo3	GGTAAAACGGGTATTGCACTTG
PGo3N	GATCCTCTCCTTCTAGACTGGGTC
PGo4	GCATCAAAGACATTTTATGACCCG
EGo2	CTATTACTGTCAACAATGTTTCATAGAAC
EGo3	GTCCTTAAGAATAGCCTTGATG
EGo2N	GCACCAATGTATTATAATGGAG
EGo2_67.11_41	GACTTGCCGTCTTCAAATGC
FGo2	CAATCTCAAAATGCTGTAAATGTTC
FGo3	GTATGGATCTTACTCGATTTGAG
Pain_SEQ3	GTAACATATCACTAGTAAAATTTG
Pain_SEQ4	CCCTGAGAAACACCTCTTGAC
Pain_SEQ5	CCGGTGACACTGATGATTATGTAC
Pain_SEQ6	CAAGGAGGAAGAAGCAGTCATAAC
Pain_SEQ7	GGTCCCAAAATGCCTCTGC
Pain_SEQ12	GAAAGCTCCATGAAATCTAATGTC
Pain_SEQ11	CTTATCCCGGTACTAATTATTAATC
Pain_SEQ13	GATCTCGCTCGCCTCCCTC
25_SEQ3	GTAAACAATTGGTTCAATGGCC
25.12_CheckG	GTATGGATCTTACTCGATTTGAG
A60_For	CCATTGTACCACAAGGGATG
A60_Rev	CCCACATACCCGTACCCG
A60_SEQ3	GAGACCCGACTACTGCTTGG
A60_2F1	CCCATTACACATTTCCTCCCG
A60_2SEQ4	GGTTCAATTGAGCTACTCCATG
34.1_For	GCATGCGGTTCCGGGTAC
34.1_Rev	CTGCAGCTGAGTCAACATGGAG
40.58_For	GAAGCCTCATTTGAAGTGGAC
40.58_Rev1	GCAACTCCCGGAGCCACTG
40.58_Rev2	CTCATCGAATGTTTTTCACCG
51.81_For	GGGACCATTGTTGTGCGTTG
36.1_SEQ3R	CCAGTATGAGGAGATGAATGAAG
36.1_3Nr4-F1	GCTCTTCTCTTTGGGGTTTAG
37.17_revG	CCTGTAACCTCAACCTTTTCACC
47.17_Rev	GTCCTTGATACGGCATGGC
51.5_T-Rev	CTTCGTTGTTTCTTCGGGTC
35.83_For	CAGAGGCTCCGGGAGTTGC
F_SEQ3	GGACTGATATGTATGCACAAGATG
P54_SEQ3	GCTATTGGATGAAGTGCTGC
CD111_SEQ2	GTAAATGGATCAAGCCCGATAAC
CD141_SEQ2	GAGGAATCATAGGGGAAAGG
S21_For	CCGAGGCCGAGGCTACCTTAAAGAAGC
S21_Rev	CCCCATAACCTTTTAGCACCGCCTACC

Table 2.1.4.: Primers used for pyrosequencing analysis of the *Pain-1* locus on chromosome III.

Name	Primer sequence in 5'-3' orientation
Pyro Pain F	GGGACCATTTGGTGTCTGTTG
Pyro Pain RB	(Btm)GCAAAGCTCTCCACAATTGAG
Pyro PainTher F	GCCTTTTGCCATGGTTCCTG
Pyro PainTherRB	(Btm)GCAGTGGTCGGGTCTCTAAAGTC
Pyro PainP40 RB	(Btm)CTGAGTCAACATGGAGTAGC
Pyro PainP40 F	GGATTGGGGAAACTGATAG
Pyro PainP40 Seq	CAGTGCTTTACGACAAGAAG
Pyro PainTheSeq	CCGGTGACACTGATGATTATG
Pyro PainDiaSq1	GAGCTAACGCCAGTTTACTTC
Pyro PainDiaSq2	CATTTCTAAAGGAGCTGATGG
51.83 PyroF	CAAAATCTTGCGTACCCCAAC
51.83 PyroRevBi	(Btm)GTTTCATAAACAAGTGCAATACCCG
51.89 PyroF	CAGCTGCTCTACTAGTGGAGG
51.89 PyroRevBi	(Btm)GAGTCTCAGCTCGGCCATC
51.814 PyroF	CCTCAGAGGCTCCGGG
51.814 PyroRevBi	(Btm)CAATAATCTCATCGAATGTTTTTC
Seq_51.814Pyro	GGGAGITGCTAAACAAGTT

Table 2.1.5: Primers used for pyrosequencing analysis of the gene *invGE* on chromosome IX.

Name	Primer sequence in 5'-3' orientation
invGE-F-fulgth	CAATATCTTTCTCCAAAATCAC
PyroE ParentsRB	(Btm)GCCCCAAACAATATTGCCC
PyroE PSNP58Seq	GGGTTTAGAAATTTATTTATTTTG
PyroE PSNP108Se	GGGGTGTTTGCTTCTCATAATA
PyroE SatF	GCAAGGGAGAAATGTTTGAAG
PyroE SatRBio	(Btm)GTCTCATGGTAGATCTTCTAGCATC
PyroE Sat6fSeq	AAGGAATCTCAGCATCACAG
PyroE Sat4Seq	CGACTATCCAAGGTGGGCT
PyroE Sat3Seq	GCTTGICCCATTTGGGCTT
PyroE Ther F	GGTTCTCATGTGCTCAGATG
PyroE TherRBio	(Btm)GTGCATCTTAGGTACATCCATG
PyroE Ther1Seq	CTGAGACAATCACAATTGAGAC
PyroE Ther6fSeq	GTAGCTGAGAGTTTTGGTGCTG
PyroE Ther2Seq	GGATATGTAGATGTAGATTTAGTAGAC
PyroE Dia F	CCTTAAGAATAGCCTTGATGTT
PyroE Dia RBio	(Btm)CTCCCTTGCTCAACTTCTTG
PyroE Dia3Seq	CCTGATAACAATTCTATCGATG
PyroE Dia2Seq	CTCGAICITAGTGGTAAACA
PyroE Dia6fSeq	GAATTCAAGCTATTCCGC

Table 2.1.6: Primers used for pyrosequencing analysis of the gene *invGF* on chromosome IX.

Name	Primer sequence in 5'-3' orientation
PyroF_4d_F	GATCCAATGTTTTCTGGCTG
PyroF_4d-RBio	(Btm)GATCCAATGTTTTCTGGCTG
PyroF_Sat_F	GGACTTAATCAATTGGATCAAT
PyroF_Sat_RBio	(Btm)CAGGCTTGATCCAATCACG
PyroF_Sat2Seq	GATCAATTIGGTACITGGTCTG
PyroF_Sat3Seq	GCIAACCAAACCTCAAGTTCAAA
PyroF_SNP58Seq	CCAGTTATCTTAGTTTGCTTTTT
PyroF_SNP96Seq	CAATAATGTTGTTTTTGCTTCTC
PyroF_SNP111Seq	GCTTCTCAIAAAGTTTTTATT
PyroF_P40/54_F	GACTCAAGGGTGCAGATGTACAAG
PyroF_P40/54_RB	(Btm)CCTCGATGTTATACATGTCTTTCC
PyroF_P40Seq	CTGGATTTGTGGATGTTGATTT
PyroF_P54Seq	GCACAACAAAATTACAAGGTTC

Table 2.1.7: Primers used for expression analysis of the gene *invGF* on chromosome IX.

Name	Primer sequence in 5'-3' orientation
F1_F	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAG
F_Expr_Rev	GTTACCAGGTAGAAATAGTTGC

Table 2.1.8: Primers used for association analysis.

Name	Primer sequence in 5'-3' orientation
pCD141_3F	ACAAGTTTGGATAAGGCAGAG
pCD141_3R	TAGAGTCTCAATTGTGATTCTCTCC

Table 2.1.9: Primers used for quantification of real-time PCR products in SYBR® Green based detection assays.

Name	Primer sequence in 5'-3' orientation
Pyro_Pain_F	GGGACCATTTGGTGTCTGTTG
Pyro_Pain_R	GCAAAGCTCTCCACAATTGAG
EF1 α _F (NICOT ET AL., 2005)	ATTGGAAACGGATATGCTCCA
EF1 α _R	TCCTTACCTGAACGCCTGTCA
L2_For (NICOT ET AL., 2005)	GGCGAAATGGGTCTGTGTTAT
L2_Rev	CATTCTCTCGCCGAAATCG

Table 2.1.10: Primers used for BAC analysis and sequencing, colony-lift, and Southern blot analysis.

Name	Primer sequence in 5'-3' orientation
<i>A) Primers for probes:</i>	
Pain1-5f	CGGAATTGGATTGTGGAATTG
Pain1-5r	TGGCGTTAGCTCAGATAGCTT
Pain_SondF1	CAATACAATCCAGATTCAGCTATTTG
Pain_SondR	CTGCAGCTGAGTCAACATGGAGTAG
CD111S2_For	CCTAGCAAAAATCGAAGAATTATGTG
CD111S1_Rev	CTTGTACATTGTCTTATCATTCTTG
pCD141_3F	ACAAGTTTGGATAAGGCAGAG
pCD141_3R	TAGAGTCTCAATTGTGATTCTCTCC
<i>B) Primers for BAC sequencing</i>	
37_F1	GCTGATTGGTTAGGTTGACTG
37_R1	CATACCGTTAATCCAGTTTTTAG
37_F2 NEW	CAGATTGATCATTTCGGTAGTGG
37_R2	GTAAAGATCAGCCCAACTAGG
37_F3	GCCAAAATATATGACGTAAAGATG
37_R3	CCAATTTCCAGTACCGGG
37_F4	CAAACATCCCTCAAGCCTAAG
37_R4	GGAACAATTGAGATTTGATTAGTC
37_R4N	GAATCACCATTGATCTGTTG
37_F5	GAGGAATAAAGAAAAGTCAAAATG
37_F6	CACTTTTCAATGATGATAACGACG
37_F7	CATGAGTTGGTTCCCATTAGAAAC
37_F8	GTTATTCATAGACACAAGAAAAAGC
163_R1	CCTTGAGGCATCGGAACAC
146_F1	CCTATGCATACTACACTTCGATC
146_F4	
146_R3	
146_F5	
239_Seq7	GTCCACTTCAAATGAGGCTTC
P40_BACcheck_R	GTTTCATCCAATTTTTTTTGAGG
Seq8	GATGGATTGGGGAAACTGATAG
Seq9	CCGTAAAATCATCAAATCTATG
Seq10	GTCCATACCTGTACAGATGCCC
BAC_CD111_For	GAATTGGTCATTTGGTCACTCTTCTTATTG
BAC_CD111_Rev	GTGTGAAGCATTAAACGCCATTGTTGAAAG
BAC_CD141_For	CAACAATGGCGCGGAGAGAATCACAATTG
BAC_CD141_Rev	CTAACATCAACATGGCTAGTAGTAGATTGC

2.1.5 Plasmids

Plasmids used for molecular cloning of cDNA and genomic DNA fragments are listed below.

Table 2.1.11: List of used vectors.

Vector	Supplied/provided by	Resistance
pGEM [®] -T Vector System	Promega	amp ^R
pGEM [®] -T Easy Vector System	Promega	amp ^R
112 A1 NE	Lothar Willmitzer, MPI for Molecular Plant Physiology/Golm	amp ^R

2.1.6 Bacterial strains

For standard cloning electrocompetent or chemical competent cells (Invitrogen, Karlsruhe) of *Escherichia coli* (*E. coli*) strains DH5 α and DH10B were used (HANAHAN, 1983).

MAX Efficiency[®] DH5 α [™] Competent cells:

F- ϕ 80*lacZ* Δ M15 Δ (*lacZYA-argF*) U169 *recA1 endA1 hsdR17* (*r_k*⁻, *m_k*⁺) *phoA supE44* λ -*thi*⁻1 *gyrA96 relA1*

ElectroMAX[™] DH10B[™] Cells:

F- *mcrA* Δ (*mrr-hsdRMS-mcrBC*) ϕ 80*lacZ* Δ M15 Δ *lacX74* *recA1 endA1 araD139* Δ (*ara, leu*)7697 *galU galK* λ -*rpsL nupG*

2.1.7 Yeast strains

In this work the yeast invertase mutant strain *SUC2* (GOZALBO & HOHMANN, 1989), and as a wild type reference the yeast strain *FY 1679* were used. Both strains were obtained from EUROSCARF: European *Saccharomyces Cerevisiae* Archives for Functional Analysis, Frankfurt.

SUC2=YIL162w: BY4741; Mat a; his3D1; leu2D0; met15D0; ura3D0; YIL162w::kanMX4

FY1679: MATa/MAT α ; ura3-52; trp1 Δ 63/TRP1; leu2 Δ 1/LEU2; his3 Δ 200/HIS3; GAL2/GAL2

2.1.8 Plant Material

Potato (*Solanum tuberosum* L.) plants used in this study were obtained from the companies SAKA-RAGIS Pflanzenzucht (SAR), Böhm-Nordkartoffel Agrarproduktion (BNA), and NORIKA (NOR). The genotypes resulted from a number of crosses among different varieties and breeding clones and were selected to represent the variation at the invertase loci, which were the objects of interest in this study. The following tetraploid cultivars were used: ‘Satina’ (SAR), ‘Diana’ (SAR), ‘Theresa’ (BNA), ‘Saturna’ (BNA), and ‘Desireé’.

Furthermore, the diploid potato genotypes P18, P40, and P54 were included because they were used as parents for mapping QTLs and candidate genes for cold sweetening of potato tubers (MENÉNDEZ ET AL., 2002).

2.2 Methods

2.2.1 Plant work

Potato tubers were germinated after the tuber dormancy period (storage at 4°C) of ca. 3 months. In some cases tuber dormancy were broken by the application of a solution containing 10g thio-urea and 2mg giberelline in 1l H₂O. Tubers were cut at the navel and incubated in the solution for 20min at RT. Afterwards the tubers were dried over night, planted in pots containing a mixture of peat and soil, and placed in a warm surrounding.

The plants were either grown in greenhouse (temperature day: 20-24°C; temperature night: 18°C; light: 6-21:00 o'clock) or in Saran-houses under natural occurring conditions through May to September. Leaf and flower material were taken during all stages, whereas tubers were only selected after natural total tuber ripening. For further analysis tubers were stored at 4°C in dark environment.

2.2.2 Bacterial work

E. coli was incubated at 37°C with LB media over night (SAMBROOK & RUSSELL, 2001). Plate incubation was performed with LB media + 1% agar, liquid cultures were incubated on a shaker (200rpm).

Transformation of constructs was performed as described by HANAHAN (1983).

2.2.3 Yeast work

The yeast strains used in this study were grown on media with a suitable carbohydrate source (2%) depending on the genotype of the strain. The invertase mutant strain *SUC2* was grown on YPD media at 30°C overnight. The wild type strain *FY 1679* was either grown on YPD media or selective media (SD) with 2% sucrose as carbohydrate source. Transformed *SUC2* yeast strains were only grown on selective SD media with sucrose as carbohydrate source. Plate incubation was performed with yeast media + 2% agar, liquid cultures were incubated on a shaker (200rpm).

For functional complementation the invertase mutant strain *SUC2* was used. Yeast transformation was performed using a simplified method described by GIETZ & SCHIESTL (1995). The strain was grown in YPD media as a 5ml overnight culture. The next day a 50ml flask with YPD was inoculated with 1ml of the overnight culture and shaken for 3-5h at 30°C. Afterwards the culture was centrifuged at 4000rpm for 5min at RT and the pellet was washed

in 25ml H₂O and centrifuged again. The washed pellet was resuspended in 1ml 100mM LiAc, transferred to a 1,5ml tube and centrifuged at 13,000rpm for 10sec. After discarding the supernatant the pellet was resuspended in 500µl 100mM LiAc.

The transformation reaction contained following components.

Table 2.2.1.: Components of the transformation reaction.

Component	Volume for one transformation
PEG 3000 (50%)	240µl
1M LiAc (pH 8.5)	35µl
ss-herring sperm (2mg/ml, denaturated at 100°C and kept on ice)	50µl
<i>SUC2</i> yeast mutant	50µl

The concentration of the plasmid used for transformation varied between 0.8-1.5µg. 350µl of the transformation reaction were added to the plasmid (in 50µl), mixed and incubated for 30min at 30°C. Afterwards the reaction was incubated for 25-30min at 42°C and centrifuged at 13,000 for 10sec. The pellet was resuspended either in 150µl 1M sorbitol or in 150µl H₂O, plated on selective SD plates and incubated at 30°C between 3 and 5d. Transformants were transferred to fresh selective SD plates and check on their inserts using colony PCR.

2.2.4 Molecular biological methods

If not indicated otherwise, the methods applied in this study were taken from SAMBROOK & RUSSELL (2001) using standard procedures.

2.2.4.1 Genomic DNA extraction from leaf tissue

Young, healthy potato leaves were harvested and immediately frozen in liquid nitrogen. Frozen samples were freeze-dried (Eps1-15, Typ 1815, Christ Gefriertrocknung GmbH, Osterode) and stored in air-tight containers at -20°C. Total genomic DNA was extracted from 0.3-0.4g freeze dried leave material according to BORMANN ET AL. (2004).

2.2.4.2 Plasmid DNA isolation from bacteria

Small scale and midi scale plasmid isolation from *E. coli* were performed using the column-based Plasmid Isolation Mini or Midi Kit (Qiagen, Hilden) according to the supplier's protocol.

Isolation of BAC DNA has been adapted by customers from the Qiagen® Plasmid Midi Kit protocol. The procedure has been used to isolate 50-200kb BAC DNA of two potato-BAC

libraries BA and BC (BALLVORA ET AL., 2002, 2007). The yield of BAC DNA from a 100ml culture was up to 25µg.

2.2.4.3 Separation of DNA fragments by agarose gel electrophoresis

DNA fragments were mixed with DNA loading buffer and analyzed by agarose gel electrophoresis. The agarose concentration depended on the size of the fragments to be resolved. Electrophoresis was performed at 5V/cm using TAE buffer. Different DNA size markers were used to estimate the size of DNA fragments. The Fermentas ladders 100bp plus and 1kb were used. After electrophoresis, DNA was visualized on a transilluminator under UV light (254nm).

2.2.4.4 Separation of DNA fragments using pulse field gel electrophoresis (PFGE)

Pulse field gel electrophoresis (PFGE) was used to estimate the size of BAC inserts. 20µg of BAC DNA in a 25µl reaction was digested with NotI and incubated for 4h at 37°C. After inactivation of the reaction at 65°C for 15min, 5µl of the reaction were loaded on a 1% agarose gel to check the restriction reaction.

Preparation of the PFGE gel:

A 1% PFGE gel was prepared using Gold Agarose and 0.5x TAE buffer. After loading 20µl of the samples and the Fermentas markers lambda ladder and mid range, the slots were sealed with liquid gel and put in the PFGE chamber.

Running of the PFGE gel:

2.5l of 0.5x TAE buffer was filled in the PFGE chamber and cooled for 20min to 11-14°C.

The machine settings were:

initial time	5sec
final	15sec
run	14h
voltage	5V/cm
incl. angle	120
pump	80

After the run, the PFGE gel was stained in a 2% EtBr solution for 1h. BAC DNA fragments were visualized on a transilluminator under UV light (254nm).

2.2.4.5 Purification of PCR products and gel-extracted DNA fragments

PCR products, gel-extracted PCR products and restriction digested products were either purified using the Roche High Pure PCR Purification Kit or using the Qiagen PCR Purification Kit following the supplier's protocol. PCR products for sequencing were purified using ExoSAP-IT[®].

2.2.4.6 Total RNA extraction from leaf and floral tissue

Total RNA from leaves and inflorescences were extracted using the ToTally RNA Isolation Kit from Ambion following the manufacturer's protocol. Additionally, the RNA was purified from possible genomic DNA contaminations using the DNA-free[™] Kit purchased from Ambion. The concentration and quality of the RNA was determined by measuring the ratios of absorbance $A_{260\text{nm}}/A_{280\text{nm}}$ which should be between 1.8 and 2.0 and $A_{260\text{nm}}/A_{230\text{nm}}$ which should be between 2 and 3 using the Nanodrop ND-1000 spectrophotometer (Peclab, Erlangen). The integrity of the RNA was tested by running a 1% agarose gel which were loaded with 300-500ng of total RNA that was mixed with 5 μ l of DNA loading dye. Total RNA was stored at -80°C .

2.2.4.7 Total RNA extraction from tuber tissue

Total RNA from tubers was extracted using the Plant RNA Isolation Kit from Invitrogen following the manufacturer's protocol. Small amounts of tubers were grinded in liquid nitrogen using mortar and pestle. Larger amounts of tuber tissue were crushed in liquid nitrogen cooled cylinders using the Retsch[®] Schwingmühle MM400 (Retsch GmbH, Haan) following the manufacturer's protocol.

Total RNA from tubers was further purified through additional precipitation steps: a high salt precipitation to remove polysaccharides and a lithium chloride precipitation to remove high molecular weight RNA. The RNA solution was adjusted to a total volume of 1ml by adding RNase free water. 250 μ l of isopropanol and 250 μ l of high salt solution (1.2M sodium citrate, 0.8M sodium chloride) were added, mixed and incubated on ice for 2h. The RNA was recovered by centrifugation at 13,000rpm for 30min at 4°C . The supernatant was removed and the pellet was rinsed with 70% ethanol, and centrifuged at 13,000rpm for 5min at 4°C . After removing the ethanol, the pellet was air-dried and dissolved in an appropriate volume of RNase free water to achieve a minimum concentration of 200ng/ μ l of total RNA.

The precipitation of high molecular weight RNA was done using 0.5 volume of 5M LiCl. The solution was mixed and kept on ice over night in the 4°C room. The RNA was recovered by

centrifugation at 13,000rpm for 30min at 4°C. After removal of the supernatant, the pellet was rinsed with 70% ethanol and centrifuged again at 13000rpm for 5min at 4°C. The ethanol was removed, the pellet air-dried and dissolved with RNase free water to a final volume of 20 to 50µl depending on the pellet size. Additionally, the RNA was purified from possible genomic DNA contaminations using the DNA-freeTM Kit purchased from Ambion. The concentration and quality of the RNA was determined by measuring the ratios of absorbance A_{260nm}/A_{280nm} which should be between 1.8 and 2.0 and A_{260nm}/A_{230nm} which should be between 2 and 3 using the Nanodrop ND-1000 spectrophotometer (Peclab, Erlangen). The integrity of the RNA was tested by running a 1% agarose gel which were loaded with 300-500ng of total RNA that was mixed with 5µl of DNA loading dye. Total RNA was stored at -80°C.

2.2.4.8 cDNA synthesis

For first strand cDNA synthesis 2.0µg of total RNA was used to synthesize cDNA by reverse transcription using 200U of SuperscriptTM II reverse Transcriptase (Invitrogen, Karlsruhe) per reaction and oligo(dT)₁₆₋₁₈ primers (500ng), (Roche, Mannheim) as priming method. cDNA synthesis was performed according to the manufacturer's protocol and additional treated with RNase H (Roche, Mannheim) for 20min at 37°C.

Synthesized first strand cDNA was either used for molecular cloning (1µl per reaction) or in qRT-PCR (2.5µl of a 1:100 dilution).

2.2.4.9 Standard Polymerase Chain Reaction (PCR)

PCR reactions were performed in different thermal cyclers. The following machines were used.

Table 2.2.2: PCR machines.

Machine	Provided by
MJ Research DNA Engine Tetrad [®] 4 peltier thermal cycler	Biozym, Hess. Oldendorf
Biometra [®] T1-Thermocycler	Biomedizinische Analytik GmbH, Göttingen
Biometra [®] T3-Thermocycler	Biomedizinische Analytik GmbH, Göttingen
Labcycler	Sensoquest Biomedizinische Elektronik, Göttingen

For standard reactions and colony-PCR *Thermus aquaticus* DNA Polymerase (*Taq*-DNA Polymerase) was used. *Taq*-DNA Polymerase was prepared described by PLUTHERO (1993). For high accuracy PCR reaction (cloning), the proof-reading Roche Fast Start High Fidelity PCR System and the proof-reading KOD Hot Start DNA Polymerase (Novagen, Darmstadt) were used.

Standard PCR reactions were performed as follows:

Table 2.2.3: Standard PCR reaction using *Taq*-DNA Polymerase

Concentration	Reagent
50-200ng	Template (genomic DNA, plasmid DNA, cDNA, bacterial or yeast colony)
2.5µl	10x PCR buffer (without MgCl ₂)
3mM	MgCl ₂
25mM	dNTP
10-25pmol	Primer 1
10-25pmol	Primer 2
0.5-0.7µl	<i>Taq</i> -DNA Polymerase (0.025-0.05U/µl)
adjust to 25µl	

Before using bacterial colonies for PCR, colonies were picked from the plate, resuspended in 15µl H₂O and denaturated at 95°C for 10min. Afterwards 1µl of the denaturated colony was used for PCR.

Before using yeast colonies for PCR, fresh grown colonies were picked and resuspended in 15µl 2mM NaOH and denaturated at 95°C for 10min. Afterwards 1µl of the denaturated colony was used for PCR.

PCR program:	Initial denaturation	94°C for 3min
	Denaturation	94°C for 30sec
	Annealing	55-65°C for 30sec
	Elongation	1min/1kb
	Final extension	72°C for 10min
	Number of cycles	30-50

Table 2.2.4: Standard PCR reaction using Roche Fast Start High Fidelity PCR System

Concentration	Reagent
50-200ng	Template (genomic DNA, plasmid DNA, cDNA)
2.5µl	10x PCR buffer (without MgCl ₂)
3mM	MgCl ₂
25mM	dNTP
10-25pmol	Primer 1
10-25pmol	Primer 2
0.25µl	Fast Start High Fidelity Polymerase (2.5U)
adjust to 25µl	

PCR program:	Initial denaturation and activation	94°C for 2min	}	10x
	Denaturation	94°C for 15sec		
	Annealing	55-65°C for 30sec		
	Elongation	1min/1kb	}	20x
	Denaturation	94°C for 15sec		
	Annealing	55-65°C for 30sec		
	Elongation	1min20sec/1kb		
	Final extension	68°C for 10min		
Number of cycles		30		

Table 2.2.5: Standard PCR reaction using Novagen KOD Hot Start DNA Polymerase

Concentration	Reagent
50-200ng	Template (genomic DNA, plasmid DNA, cDNA)
2.5µl	10x PCR buffer (without MgCl ₂)
1mM	MgSO ₄
25mM	dNTP
10-25pmol	Primer 1
10-25pmol	Primer 2
0.5µl	KOD Hot Start DNA Polymerase(0.5U)
adjust to 25µl	

PCR program:	Initial denaturation and activation	94°C for 2min
	Denaturation	94°C for 15sec
	Annealing	55-65°C for 30sec
	Elongation	1min/1kb
	Number of cycles	30-50

2.2.4.10 Molecular cloning of cDNA constructs for yeast *SUC2* transformation

Unless otherwise described, cloning strategies performed in this study included directional cloning of PCR products into pGEM[®]-T Vector System or pGEM[®]-T Easy Vector System (Promega, Mannheim). All reactions were performed following the manufacturer's protocol. All constructs were verified by sequencing. Alleles were defined as sequences, which were found twice in two independent PCRs. In some cases where sequences varied a lot, the allelic consensus sequence was used for allele definition.

Production of the final expression vector constructs for yeast transformation was performed using the 112 A1 NE yeast expression vector (Appendix A 2) kindly provided by Lothar Willmitzer's Group, MPI for Molecular Plant Physiology/Golm. 112 A1 NE inserts are driven under the control of the constitutive strongly expressed *Adh1* promoter. The vector, originally

an *E. coli* vector, also includes the *AdhI* terminator, an ampicillin resistance for *E. coli* selection, and tryptophane for yeast selection. pGEM[®]-T constructs bearing invertase alleles were amplified using proof-reading high fidelity *Taq*-Polymerase (Roche, Mannheim) to insert specific cloning sites. Alleles of the loci *Pain-I* and *invGF* were amplified with primers inserting *NotI* and *BamHI* restriction sites for sticky end ligation into the vector 112 A1 NE. *Pain-I* and *invGF* PCR products as well as the vector were digested with the corresponding restriction enzymes, followed by ligation of the linearized vector with the digested PCR products. Alleles of the *invGE* locus were amplified with primers inserting only *NotI* restriction sites because a *BamHI* restriction site is included in the gene. *invGE* PCR products as well as the vector were digested with *NotI* restriction enzyme, followed by blunt end ligation of the linearized vector with the digested PCR product. 112 A1 NE constructs were verified by sequencing.

All constructs generated in this study are indicated in the Table below.

Table. 2.2.6: Overview of generated constructs.

Gene	Vector	Primer pairs used (Table 2.1.2)
<i>Initial cloning of invertase alleles:</i>		
<i>Pain-I</i>	pGEM [®] -T/ T Easy	ZrPain-F/ PainUni-R
<i>invGE</i>	pGEM [®] -T/ T Easy	invGE-F-fulgth/ invGE-R-fulgth
<i>invGF</i>	pGEM [®] -T/ T Easy	invGF-F-fulgth/ invGF-R-fulgth
<i>pCD111</i>	pGEM [®] -T/ T Easy	CD111fl_F/ CD111fl_R
<i>pCD141</i>	pGEM [®] -T/ T Easy	CD141fl_F/ CD141fl_R
<i>Cloning of constructs for yeast transformation:</i>		
<i>Pain-I</i>	112 A1 NE	PainNotI_R/ PainBamHI_R
<i>invGE</i>	112 A1 NE	ENotI_F/ ENotI_R
<i>invGF</i>	112 A1 NE	FNotI_F/ FBamHI_R

The presence of the transformed plasmid was tested via colony-PCR using a primer that was located in the backbone of the vector and a second primer localized within the insert sequence (Table 2.1.3, primer pair T7/SP6P).

2.2.4.11 Quantitative real-time PCR

Quantitative real-time PCR (qRT-PCR) was performed using a Mastercycler[®] ep *realplex* (Eppendorf, Hamburg). Primers were tested for undesired primer dimer formation by melting curve analysis (55°C to 95°C with a heating rate of 0,1°Cs⁻¹ and continuous fluorescence measurement) using the Power SYBR[®] Green PCR Master Mix (Applied Biosystems,). For standard curves tuber cDNA of the 4th week of cold storage were diluted 1:10, 1:100 and 1:1000. Total invertase transcripts were quantified using the primer pair Pyro_Pain_F/Pyro_Pain_R in a PCR with a 100 fold template dilution. The quantification of

the normalization gene *efl- α* (elongation factor 1- α , NICOT ET AL, 2005) was calculated based on a 100 fold dilution series of the samples also tested with *Pain-1* transcript primer (qRT-primer are listed in Table 2.1.9). Additionally, the normalization gene *L2* (cytoplasmic ribosomal protein, NICOT ET AL, 2005) were used for samples of the cultivar ‘Theresa’.

Determined *L2* values were similar to values of the gene *efl- α* , and in the course of the work *efl- α* was used for subsequent qRT-PCR.

Plant material for quantitative expression analysis was grown in three independent biological replicates, from which two were used for all analysed samples. In case of high variances within the measured values, the third replicate was also tested. qRT-PCR reactions of each of the biological replicates were done in technical duplicates.

Table 2.2.7: Reaction setup for qRT-PCR

Concentration	Reagent	Volume
1:10, 1:100, 1:1000	cDNA	2.5 μ l
2x	Power SYBR [®] Green PCR Master Mix	12.5 μ l
10pmol	Primer 1	2 μ l
10pmol	Primer 2	2 μ l
fill to 25 μ l with H ₂ O		

PCR program for qRT-PCR of invertase transcripts:

Initial denaturation	95°C for 10min
Denaturation	95°C for 15sec
Annealing	55°C for 30sec
Elongation	72°C for 45sec
melting curve analysis	
Number of cycles	50

PCR program for qRT-PCR of the normalization gene *EF1 α* :

Initial denaturation	95°C for 10min
Denaturation	95°C for 15sec
Annealing	60°C for 30sec
Elongation	72°C for 45sec
melting curve analysis	
Number of cycles	40-50

2.2.4.12 Sequencing

DNA sequencing was performed by the MPIZ DNA core facility on Applied Biosystems (Weiterstadt) Abi Prism 377, 3100 and 3730 sequencers using BigDye-terminator v3.1. chemistry. Premixed reagents were purchased from Applied Biosystems. Sequences were analysed with the software Chromas Pro version 1.33 (share-it, Cologne) or the Lasergene software.

2.2.4.13 Pyrosequencing

Pyrosequencing is a robust and quantitative method of DNA sequencing based on real-time detection of pyrophosphate, which is released as a result of nucleotide incorporation in a the ‘sequencing by synthesis’ reaction (RONAGHI ET AL., 1996). ‘Sequencing by synthesis’ involves taking a single strand of the DNA to be sequenced and then synthesizing its complementary strand enzymatically. The pyrosequencing method is based on detecting the activity of DNA polymerase with another chemiluminescent enzyme. Essentially, the method allows sequencing of a single strand of DNA by synthesizing the complementary strand along it, one base pair at a time, and detecting which base was actually added at each step. The template DNA is immobilized, and solutions of A, C, G, and T nucleotides are added and removed after the reaction, sequentially. Light is produced only when the nucleotide solution complements the first unpaired base of the template. The sequence of solutions, which produce chemiluminescent signals, allows the determination of the sequence of the template. The templates for pyrosequencing can be made both by solid phase template preparation (Streptavidin coated magnetic beads) and enzymatic template preparation (Apyrase+Exonuclease). In this study, Streptavidin coated magnetic beads were used.

PCR conditions:

The primers used for pyrosequencing analysis are listed in the Tables 2.4.1 for the *PainI* locus on chromosome III, 2.4.2 for the *invGE* locus on chromosome IX and 2.4.3 for the *invGF* locus on chromosome IX.

PCR setup and conditions are listed in Table 2.10.

Solid phase template preparation for single-stranded DNA template:**Table 2.2.8: Reaction setup for pyrosequencing**

Volume	Reagent
20µl	PCR product
15µ	H ₂ O (LiCroSolv Merck)
40µl	Binding buffer (10mM Tris-HCl, 2M NaCl, 1mM EDTA, 0.1% Tween 20; pH 7.6)
5µl	Streptavidin-coated Super Paramagnetic beads

The reaction mixture was incubated for 5min on a shaker at RT. The Streptavidin-template complex was captured using the PSQ96 Sample Prep tool (Biotage AB, Uppsala, Sweden) and single-stranded template was generated by washing in 70% EtOH for 3sec, in 0.2M NaOH for 5sec followed by washing in washing buffer (10mM Tris acetate pH 7.6) for 10sec. 1µl of the sequencing primer (0.25mM) was annealed to the immobilized template in 39µl annealing buffer (20mM Tris acetate, 2mM Mg acetate, pH 7.6) and heated at 80°C for 2min followed by slow cooling to room temperature.

Pyrosequencing:

The sequencing reaction was performed automatically with the PSQ 96 system (Biotage AB, Uppsala, Sweden) at 28°C using a SNP reagent kit according to the manufacturer's protocol. The ssDNA template is hybridized to a sequencing primer and incubated with the enzymes DNA polymerase, ATP sulfurylase, luciferase and apyrase, and with the substrates adenosine 5' phosphosulfate (APS) and luciferin. The four enzyme mixture, the substrates and the four separate deoxynucleotide triphosphates were loaded into the reagent cartridge (PSQ 96 SNP Reagent Kit, Biotage AB, Uppsala, Sweden).

1. The addition of one of the four deoxynucleotide triphosphates initiates the second step. DNA polymerase incorporates the correct, complementary dNTPs onto the template. This incorporation releases pyrophosphate (PPi) stoichiometrically.
2. ATP sulfurylase quantitatively converts PPi to ATP in the presence of adenosine 5' phosphosulfate. This ATP acts as fuel to the luciferase-mediated conversion of luciferin to oxyluciferin that generates visible light in amounts that are proportional to the amount of ATP. The light produced in the luciferase-catalyzed reaction is detected by a camera and analyzed in a program.
3. Unincorporated nucleotides and ATP are degraded by the apyrase, and the reaction can restart with another nucleotide.

The following Tables contain information about primers used in the pyrosequencing assay for detection of more than one SNP to dissect the corresponding alleles due to the complexity of the approach. Tables for the dissection of two alleles are not shown because the setup is described in the corresponding result section.

Table 2.2.9: Primers used in the pyrosequencing assay of *invGE* alleles from the tetraploid cultivars.

Genotype	Allele	SNP position	Allele specific SNP	Sequencing primer
‘Satina’	<i>E_SA</i>	SNP 1237	<i>E_SA/E_SN1/E_SN2/E_SN3</i> T/G/G/G	PyroE_Sat6fSeq
	<i>E_SN1</i>	SNP 1366	A/G/A/A	PyroE_Sat4Seq
	<i>E_SN2</i>	SNP 1379	C/C/T/C	PyroE_Sat3Seq
	<i>E_SN3</i>	SNP 1216	A/A/A/G	PyroE_Dia_F
‘Diana’	<i>E_DA</i>	SNP 1086	<i>E_DA/E_DN1/E_DN2</i> A/T/T	PyroE_Dia6fSeq
	<i>E_DN1</i>	SNP 1117	T/C/T	PyroE_Dia2Seq
	<i>E_DN2</i>	SNP 924	T/T/C	PyroE_Dia3Seq
‘Theresa’	<i>E_TA</i>	SNP 1615	<i>E_TA/E_TN1/E_TN2/E_TN3</i> A/T/T/T	PyroE_Ther6fSeq
	<i>E_TN1</i>	SNP 1720	C/A/C/C	PyroE_Ther1Seq
	<i>E_TN2</i>	SNP 1553	T/T/C/T	PyroE_Ther2Seq
	<i>E_TN3</i>	SNP 1473	T/T/T/G	PyroE_Ther_F

For PCR fragment amplification the following primers were used (Table 2.1.5): ‘Satina’:

PyroE_SatF/PyroE_SatRBio; ‘Diana’: PyroE_Dia_F/PyroE_Dia_RBio; ‘Theresa’:

PyroE_Ther_F/PyroE_TherRBio.

Table 2.2.10: Primers used in the pyrosequencing assay of *invGE* alleles from the diploid genotypes.

Genotype	Allele	SNP position	Allele specific SNP	Sequencing primer
P18	<i>E_P18N1</i>	SNP 108	A	PyroE_PSNP108Se
	<i>E_P18N2</i>		T	
P40	<i>E_P40N1</i>	SNP 58	C	PyroE_PSNP58Seq
	<i>E_P40N2</i>		T	
P54	<i>E_P54N1</i>	SNP 58	C	PyroE_PSNP58Seq
	<i>E_P54N2</i>		T	

For PCR fragment amplification the following primers were used (Table 2.1.5): P18, P40, P54: *invGE*-F-fulgth/PyroE_ParentsRB.

Table 2.2.11: Primers used in the pyrosequencing assay of *invGF* alleles from the tetraploid cultivars.

Genotype	Allele	SNP position	Allele specific SNP	Sequencing primer
			<i>F_SN1/F_SN2/F_SN3/F_SN4</i>	
‘Satina’	<i>F_SN1</i>	SNP 111	T/C/C/C	PyroF_SNP111Seq
	<i>F_SN2</i>	SNP 459	C/T/C/C	PyroF_Sat2Seq
	<i>F_SN3</i>	SNP 378	A/A/G/A	PyroF_Sat3Seq
	<i>F_SN4</i>	SNP 96	C/C/C/T	PyroF_SNP96Seq
‘Diana’	<i>F_DN1</i>	SNP 96	T	PyroF_SNP96Seq
	<i>F_DN2</i>		C	
‘Theresa’	<i>F_TN1</i>	SNP 96	T	PyroF_SNP96Seq
	<i>F_TN2</i>		C	

For PCR fragment amplification the following primers were used (Table 2.1.6): ‘Satina’: PyroF_Sat_F/PyroF_Sat_RBio and PyroF_4d_F/PyroF_4d_RBio for the alleles *F_SN1* and *F_SN4*; ‘Diana’ and ‘Theresa’: PyroF_4d_F/PyroF_4d_RBio.

Table 2.2.12: Primers used in the pyrosequencing assay of *invGF* alleles from the diploid genotypes.

Genotype	Allele	SNP position	Allele specific SNP	Sequencing primer
P40	<i>F_P40N1</i>	SNP 1534	G	PyroF_P40Seq
	<i>F_P40N2</i>		A	
P54	<i>F_P54N1</i>	SNP 1446	T	PyroF_P54Seq
	<i>F_P54N2</i>		C	

For PCR fragment amplification the following primers were used (Table 2.1.6): P40 and P54: PyroF_P40/54_F/PyroF_P40/54_RB.

2.2.4.14 Single stranded conformation polymorphism (SSCP) analysis

SSCP is the electrophoretic separation of single-stranded nucleic acids based on subtle differences in sequence (often a single base pair), which results in a different secondary structure and a measurable difference in mobility through a gel.

Background:

The mobility of double-stranded DNA in gel electrophoresis is dependent on strand size and length but is relatively independent of the particular nucleotide sequence. The mobility of single strands, however, is noticeably affected by very small changes in sequence, possibly one changed nucleotide out of several hundred. Small changes are noticeable because of the relatively unstable nature of single-stranded DNA; in the absence of a complementary strand, the single strand may experience intrastrand base pairing, resulting in loops and folds that give the single strand a unique 3D structure, regardless of its length. A single nucleotide change could dramatically affect the strand's mobility through a gel by altering the intrastrand base pairing and its resulting 3D-conformation (MELCHER, 2003).

Single-strand conformation polymorphism analysis takes advantage of this quality of single-stranded DNA. First announced in 1989 as a new means of detecting DNA polymorphisms, or sequence variations, SSCP analysis offers an inexpensive, convenient, and sensitive method for determining genetic variation (SUNNUCKS ET AL., 2000).

Like restriction fragment length polymorphisms (RFLPs), SSCPs are allelic variants of inherited, genetic traits that can be used as genetic markers. Unlike RFLP analysis, however, SSCP analysis can detect DNA polymorphisms and mutations at multiple places in DNA fragments (ORITA ET AL., 1989). As a mutation scanning technique, though, SSCP is more often used to analyze the polymorphisms at single loci, especially when used for medical diagnoses (SUNNUCKS ET AL., 2000).

The SSCP analysis of the invertase genes *invGE* and *invGF* was published by LI ET AL. (2005). The invertase genes *Pain-1* and *pCD141* were analyzed also by LI ET AL. (2008). Primers for *pCD141* (pCD141_3F/pCD141_3R) were generated in the course of this work (Table 2.1.8).

SSCP procedure was performed as described in the cited publications.

2.2.4.15 BAC DNA library screens

BAC library screens were performed using two different libraries (supplied by LION Bioscience AG, Heidelberg). The first library, so called 'BA' library (BALLVORA ET AL., 2002), was constructed from partially *HindIII* digested high-molecular weight genomic DNA of the potato genotype P6/210 in the binary vector pCLD04541. P6/210 is a F1 hybrid of the parental clones P41 (H79.1506/1) and P40 (H80.696/4), (RITTER ET AL., 1991). The BAC library consists of approximately 100.000 clones with an average insert size of 70kb. With the size of a haploid potato genome being approximately 109 base pairs, the genome coverage was 6-7 times per haploid genome.

The second BAC library, so called 'BC' library (BALLVORA ET AL., 2007), was constructed from partially *EcoRI* digested genomic P6/210 DNA in the binary vector pBeloBAC11. The average insertion size was 80kb, corresponding to an, on average, 8-fold coverage of the potato genome.

Library screens were performed using a set of four filters carrying $\approx 100,000$ clones.

Preparation of radioactively-labeled probes and DNA hybridisation:

200ng of gel-purified or ExoSAP-IT[®] purified PCR products were diluted up to a final volume of 12µl and 2.5µl buffer A1 (0.2mM dTTP, 0.2mM dCTP, 0.2mM dGTP; Invitrogen, Karlsruhe), 2.5µl DNaseI DNA Polymerase (Invitrogen, Karlsruhe) and 3µl α^{32} pdATP were added. The reaction was incubated at 16°C for 1h. The probe was purified on a Sephadex-G50 column (Amersham Biosciences, Sweden) and heated at 95°C for 5min.

Pre-hybridisation and hybridisation were carried out in hybridisation solution (20x SSPE [3M NaCl, 0.2M NaH₂PO₄, 20mM EDTA pH 7.0], 100x Denhardtts, 10% SDS) in glass tubes (30cm x 4cm) at 65°C under continuous rotation in a hybridisation oven (Bachofen, Reutlingen). Pre-hybridisation was performed at least for 1h before hybridisation or overnight. Probe hybridisation was carried out overnight.

After hybridisation the filter was washed accordingly:

1. 50ml 2x SSPE + 0.1% SDS at 65°C for 10-15min
2. 50ml 1x SSPE + 0.1% SDS at 65°C for 10min
3. 50ml 0.2x SSPE + 0.1% SDS at 65°C for 10min

The filter was wrapped in thin plastic foil (Saran wrap) and exposed overnight either to a phosphorimager screen (Molecular Dynamics) in a cassette at RT or to film (Kodak[®] X-Omat AR Film, XAR 5, 35x43cm (Sigma-Aldrich Chemie GmbH, Steinheim) in a cassette at -70°C.

The *Pain-1* locus of BC library clones BC14 and BC17 was custom sequenced by GATC Biotech, Konstanz using primer walking as sequencing method. Full-length BAC insert sequencing was performed for the *Pain-1* clones BC2 (insert size: 75kb), BC14 (insert size: 75kb), and BC15 (insert size: 97kb), and for the BC clone BC3 (insert size: 130kb) for the genes *pCD111* and *pCD141*. Full-length BAC insert sequencing was carried out by MWG Biotech AG, Ebersberg using the 454 sequencing technique on the GS FLX system.

2.2.4.16 Colony-lift

The colony-lift is an easy method to confirm hybridisation positive BAC clones. For this purpose single colonies from hybridisation positive BAC clones were grown on selective LB plates (BA BACs: Tet; BC BACs: Cam) at 37°C overnight. To transfer the DNA, Amersham Hybond[™] – N⁺ membrane (GE Healthcare, Buckinghamshire, UK) was placed on the plate and then stripped off again. The membrane was washed accordingly:

1. once 10% SDS for 3min
2. once denaturation solution (1.5M NaCl; 0.5M NaOH) for 5min
3. twice neutralisation solution (1.5M NaCl; 0.5M Tris-HCl pH 8.0) for 5min

After washing, the membrane was air-dried (ca. 15min) and then cross linked with a UV crosslinker (Stratagene) by applying 120,000J x cm⁻² of energy. The membrane was wrapped in thin plastic foil (Saran wrap) and stored in the refrigerator at 4°C.

Preparation of radioactively-labelled probes and DNA hybridisation:

Previously described in 2.2.4.15.

2.2.4.17 Southern blot analysis

Southern analysis was performed according to SAMBROOK ET AL. (1998). 20ng of purified BAC DNA were digested to completion with *HindIII*, *SmaI* and *BamHI* for 4h.

DNA separation and blotting onto membranes:

Digested BAC DNA was subsequently mixed with loading buffer, loaded on a 1% agarose gel and separated for 5h at 70V via electrophoresis. After the run, the gel was stained in a 2% EtBr solution for 15min at RT and DNA was visualized on a transilluminator under UV light (254nm). The gel was destained in H₂O for 20min at RT.

DNA was transferred on Amersham HybondTM – N⁺ membrane for 18h at RT using 0.4M NaOH buffer. After the transfer procedure DNA was cross linked to the filter with a UV crosslinker (Stratagene) by applying 120,000J x cm⁻² of energy. The blot was wrapped in thin plastic foil (Saran wrap) and stored in the refrigerator at 4°C.

Preparation of radioactively-labelled probes and DNA hybridisation:

Previously described in 2.2.4.15.

2.2.5 Three-dimensional modelling of invertase alleles

3D-modelling was performed by Pawel Durek, MPIMP/Golm. The modelling of the allelic invertase molecular structure was based on the invertase 3D and crystal structure of cyanobacteria (ALBERTO ET AL., 2004). The models are comparative, superimposing two allelic sequences. Differences of the alleles are highlighted. The modelling was applied for associated and non associated *Pain-1* alleles of the tetraploid potato cultivars ‘Satina’ and ‘Diana’. The models include the putative sucrose binding site with the substrate sucrose. In addition to the structural visualization of amino acid exchanges, also the electrostatic potential (EP) of the molecules was mapped at pH 4.7 mimicking the vacuolar and apoplastic conditions.

All models were predicted by homology modelling applying the automated structure prediction server 3djigsaw (BATES ET AL., 2001). Subsequently, the models were prepared for continuum electrostatics calculation utilizing the PDB2PQR package (Version: 1.3.0) (DOLINSKY ET AL., 2004). For the calculation, the AMBER force field and the protonation states at pH 4.7 were used (LI H. ET AL., 2005), to reflect the proper vacuolar and apoplastic acidity. The isoelectric surfaces were computed by the adaptive Poisson-Boltzmann Solver (APBS) (BAKER ET AL., 2001) utilizing standard parameters at the temperature of 298K. The

comparison of the isoelectric surfaces revealed no significant changes upon lowering the temperature parameter. The isoelectric surfaces were visualized by PyMol (DELANO, 2002).

The allel models were superimposed by Swiss-PDB Viewer (GUEx ET AL., 1997). Differences between structures were marked by color-code as judged by the RMSs between the structures and visualized by PyMol. In Figures deep blue colour indicates highly similar protein regions, whilst red characterizes high structural differences.

2.2.6 Biochemical methods

2.2.6.1 Protein extraction from yeast

50ml of yeast cultures were grown for 72h at 30°C on a shaker (200rpm). Cells were harvested $OD_{600} \approx 2.3$ by centrifugation at 4,000rpm, for 10min, at 4°C. The pellet was washed with 25ml cold dest. H₂O and centrifuged again (4,000rpm, for 10min, at 4°C). The pellet was kept on ice and resuspended in 4ml cold protein extraction buffer (25mM Tris phosphate pH 6.7; 10mM glycerol, 0.1mM DTT, 1mM EDTA, 2% Protease Inhibitor Cocktail for general use (Sigma, St. Louis, USA). After adding half of the solution volume of acid washed glass beads (425-600 micron, Sigma-Aldrich Chemie GmbH, Taufkirchen), the tubes were vortexed for 10min in time intervals of 2min, keeping the solutions on ice in between. After this procedure a centrifugation followed at 6,000rpm for 10min at 4°C. The supernatant were loaded on PD-10 columns (GE Healthcare, Buckinghamshire, UK) equilibrated with protein extraction buffer without protease inhibitors. Proteins were eluted from the column by adding 2ml of protein extraction buffer without protease inhibitors and kept on ice.

Determination of the protein concentration using the Bradford assay:

The protein concentration was determined using the Bradford assay (BRADFORD, 1976) with Bradford dye reagent (Protein assay, Bio-Rad, Hercules, USA). 2-10µl of the protein extracts were added to 198-190µl H₂O and 800µl of Bradford dye reagent. The mixture was incubated for 10-15min at RT and extinction was measured at 595nm against a blank (200µl H₂O + 800µl dye) and bovine serum albumin as a standard.

2.2.6.2 Enzymatic assay of invertase

The enzymatic invertase assay was performed using a modified protocol of ZRENNER ET AL. (1995). The principle of the enzymatic reaction is the following:



Figure 2.2.1: Simplified scheme of the principle of invertase reaction.

For a 100µl invertase assay reaction 20µg of total yeast protein extracts were used. The reaction components were the following:

Table 2.2.13: Reaction setup for invertase assay.

Concentration	Component
20µg	total yeast protein extract
20mM	NaOAc (pH 4.7)
2.5-120mM	sucrose
fill up to 100µl with H ₂ O	

The reaction was incubated for 1h at 30°C for invertase alleles from the genes *Pain-1*, *invGE*, and *invGF*. Additionally, invertase alleles from the *Pain-1* locus were incubated for 1:30h at 4°C. After incubation 10µl of 1M NaHPO₄ (pH 7.2) were added and the reaction was stopped by heating at 95°C for 10min. Blanks had the same reaction mixture, but were heat inactivated without incubation.

It was checked that the assay was linear with time for at least 90min, and was linearly dependent on the amount of protein in the crude extract added up to 50µg.

The quantitative, enzymatic determination of glucose was performed using the enzymes hexokinase (HK) and glucose-6-phosphate dehydrogenase (G6PDH). The principle of the reaction is the following:

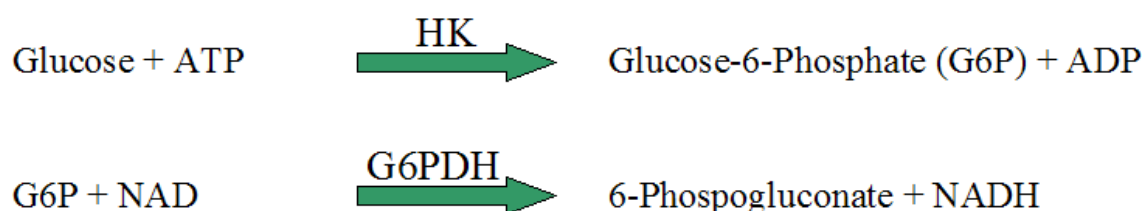


Figure 2.2.2: Simplified scheme of the NADH coupled glucose detection.

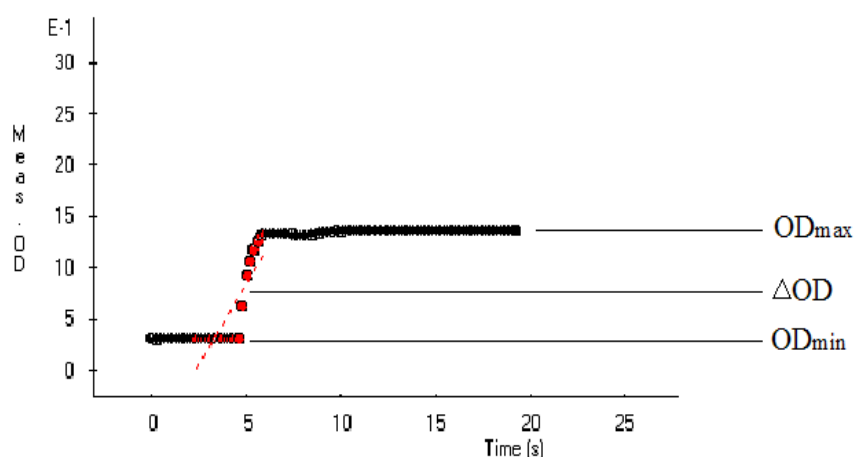
Glucose is phosphorylated by adenosine triphosphate (ATP) in the reaction catalyzed by hexokinase (HK). Glucose-6-phosphate (G6P) is then oxidized to 6-phospho-gluconate in the presence of the oxidized nicotinamide adenine dinucleotide (NAD) in a reaction catalyzed by glucose-6-phosphate dehydrogenase (G6PDH). During this oxidation, an equimolar amount of NAD is reduced to NADH. The consequent increase in absorbance at 340nm is directly proportional to glucose concentration.

The components of a 300µl reaction were the following:

Table 2.2.14: Reaction setup for glucose determination.

Stock concentration/storage	Final concentration	Volume/reaction
1M Imidazol pH 6,9 HCl / 4°C	100mM	30µl
1M MgCl ₂ / RT	5mM	1.5µl
100mM NADP+ / -20°C	2mM	6µl
100mM ATP / -20°C	1mM	3µl
G6PDH from yeast (400U/ml) / 4°C	2U	5µl
Enzymatic invertase reaction / -20°C		10µl
fill up to 294µl with H ₂ O		

G6PDH was directly resuspended in the reaction components after spinning out of ammonium sulphate. HK (300U/ml) was also spun out of ammonium sulphate and resuspended in 6µl of the reaction components without G6PDH. The reaction was started by adding HK to the assay and absorbance was measured at 340nm. For absorbance measurements, a plate reader was used and OD_{min} and OD_{max} values were determined resulting in ΔOD used for calculation



$$\text{nmol Glc/mg protein/h} = \Delta\text{OD} / 5.3 / A * B * C / D / E$$

Figure 2.2.3: Overview of plate reader output and glucose calculation. 5.3=millimolar extinction coefficient of NAD⁺, NADH, NADP⁺ or NADPH at 340nm using the 96 well plate. A=volume of invertase reaction for glucose detection, B=total volume of invertase reaction, C=total volume of glucose detection reaction, D=protein concentration used in invertase assay, E=incubation time of invertase assay.

Calculated glucose concentrations were blotted against all tested sucrose concentrations and tested for Michaelis-Menten kinetics. Transforming the measured values by Lineweaver-Burk plot, led to the determination of the Michaelis constant (K_m) and the maximal velocity (v_{max}) of invertase reaction. The K_m value describes the affinity of an enzyme to its substrate. K_m is inversely proportional to the substrate affinity, and is for a one-substrate reaction independent of the enzyme concentration. The enzymatic substrate conversion increases with increasing substrate concentration until the enzyme is saturated. This characteristic is described by the maximum velocity (v_{max}). In contrast to K_m , v_{max} depends on the enzyme concentration. Both biochemical characteristics vary with pH, with temperature and with the structure of the substrate.

Calculation of the significance values for the biochemical parameters

The significance values for differences between the K_m and v_{max} values measured for the *Pain-I* alleles at 30°C and 4°C, and for the alleles of the genes *invGE* and *invGF* at 30°C were calculated by Benjamin Stich, MPIZ/Köln.

In a first step, K_m and v_{max} estimates were calculated for each of the two biological replicates and the three technical replicates. These estimates were then used in a mixed-model context to examine:

- (i) the presence of differences between the alleles in each of the four experiments (*Pain-I* assay at 30°C and 4°C, *invGE* assay at 30°C, *invGF* assay at 30°C):

$$Y = \mu + \text{allele} + \text{allele:biolrep} + \text{error},$$

where μ was the general mean, allele an effect for the allele (without wild type reference *FY 1679*), and allele:biolrep was an effect for the biological replicate nested within the allele effect. Allele was regarded as fixed and allele:biolrep as random.

- (ii) the influence of the temperature regime (*Pain-I* assay at 30°C and 4°C):

$$Y = \mu + \text{tempreg} + \text{allele} + \text{tempreg*allele} + \text{tempreg:allele:biolrep} + \text{error},$$

where tempreg was the effect for the temperature regime, and tempreg*allele the interaction between the temperature regime and the allele. Tempreg and tempreg*allele were regarded as fixed.

- (iii) the influence of the location of the proteins (vacuolar vs. cell wall-bound):

$$Y = \mu + \text{loc} + \text{loc:allele:biolrep} + \text{error},$$

where loc was the effect for the location of the protein, which was regarded as fixed.

(iv) the presence of differences between all examined alleles:

$$Y = \mu + \text{allele} + \text{allele:biolrep} + \text{error}.$$

For fixed effects, a Wald-F-test was performed.

2.2.6.3 Protein separation by SDS-PAGE and Western blot analysis

Protein separation by SDS-PAGE:

The protein extracts were prepared for SDS-PAGE (LAEMMLI, 1970) by adding SDS sample buffer and by heating the samples at 95°C for 10min to denature the proteins.

The extracts were applied onto precast 4-12% Bis-Tris gradient gels (NuPAGE®, Invitrogen, Carlsbad, USA) and were separated in MOPS running buffer (Invitrogen, Carlsbad, USA) at 200V in XCell SureLock™ Mini-Cells (Invitrogen, Karlsruhe). The following molecular weight markers were used:

1. Spectra™ Multicolor Broad Range Protein Ladder (10-260kDa), (MBI Fermentas GmbH (St. Leon-Rot))
2. MagicMark™ XP Western Protein Standard (Invitrogen, Carlsbad, USA)

After the run the gels were stained with Coomassie Brilliant Blue (Bio-Safe™ Coomassie G-250 stain, Bio-Rad, Hercules, USA) to detect the proteins. Gels for blotting were not stained.

Western blot analysis:

The protein extracts were blotted on Amersham Hybond-P PVDF Membrane (GE Healthcare, Buckinghamshire, UK) by wet electrotransfer using a blotting module (XCell™ II Blot Module, Invitrogen, Carlsbad, USA) according to the manufacturer's instructions. After blotting, the protein transfer onto the membrane was visualized with P3504 Ponceau S (Sigma-Aldrich Chemie GmbH, Steinheim) and after destaining and TBS (50mM Tris-HCl pH 7.5, 150mM NaCl) washing, the membrane was incubated in blocking solution (TBS + 3% non-fat dried milk powder) for 1h at RT and continuously shaking. For primary antibody reaction, the membrane was incubated with polyclonal anti-invertase antibodies overnight at RT on a shaker.

Table 2.2.15: Anti-invertase antibodies used in Western blot analysis.

Antibody	Concentration	Supplied/provided by
25kDa vacuolar invertase carrot isoform I	1:400	Arnd Sturm(former member of the Friedrich Miescher Institut, Basel, Switzerland)
43kDa vacuolar invertase carrot isoform I	1:500	
vacuolar invertase carrot isoform I	1:1000	Heather A. Ross (Scottish Crop Research Institute, Dundee, Scotland)
vacuolar invertase carrot isoform II	1:1000	
58kDa vacuolar invertase potato	1:200	

All antibodies were used in TBS + 1.5% non-fat dried milk powder.

As a secondary antibody the ECLTM Anti-rabbit IgG Horseradish Peroxidase linked whole antibody (from donkey, GE Healthcare, Buckinghamshire, UK) in a concentration 1:10000 in TBS without milk powder was used and incubated for 1h at RT on a shaker. For visualisation, the Amersham ECL Plus Western Blotting Detection Reagents (GE Healthcare, Buckinghamshire, UK) was used according to the manufacturer's protocol.

3 Results

3.1 The *Pain-1* locus on chromosome III

The *Pain-1* locus encodes a soluble acid invertase, which is located in the vacuole. The pH optimum of the corresponding enzyme ranges between 4.5 and 5.0 (TYMOWSKA-LALANNE & KREIS, 1998). cDNAs of several potato soluble acid invertases have been cloned and proteins were partially characterized at functional level (ZHOU ET AL., 2004; accession L29099, ZRENNER ET AL., 1996; accession X70368, ZHANG ET AL., unpublished; accession AY341425, MATSUURA-ENDO ET AL., 2006 unpublished; accession DQ478950).

3.1.1 Structural characterization of the *Pain-1* locus

3.1.1.1 Molecular cloning of *Pain-1* invertase cDNA alleles from tuber tissue

Soluble acid invertase transcripts are strongly induced during tuber cold storage (ZRENNER ET AL., 1996; BAGNARESI ET AL., 2008). Therefore, molecular cloning of *Pain-1* alleles was performed using potato tubers stored for four weeks at 4°C in the dark.

Using the full-length *Pain-1* specific primers ZrPain-F/PainUni-R (chapter 2, Table 2.1.2), cDNA invertase alleles were cloned and sequenced from the three tetraploid potato cultivars ‘Satina’, ‘Diana’, and ‘Theresa’, and from three diploid potato genotypes P18, P40, and P54. The tetraploid genotypes are representatives for the associated SSCP fragments of the *Pain-1* gene (LI ET AL., 2008; Table 3.1.1) and, therefore, were chosen in this study. Phenotypic characterization of potato chips quality from the cultivars showed variability within the trait (Table 3.1.2). The diploid potato genotypes were included because they were used as parents for mapping QTLs and candidate genes for cold sweetening of potato tubers (MENÉNDEZ ET AL., 2002).

Table 3.1.1: Distribution of the associated SSCP fragments *Pain1-5c*, *Pain1-8c* and *Pain1-9a* present in the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’.

Genotype	<i>Pain1-5c</i>	<i>Pain1-8c</i>	<i>Pain1-9a</i>
‘Satina’	0	1	1
‘Diana’	1	1	1
‘Theresa’	0	0	0

0=SSCP fragment is absent, 1=SSCP fragment is present.

Table 3.1.2: Potato chips colour scores for the cultivars ‘Satina’, ‘Diana’, and ‘Theresa’.

Genotype	CQA	CHS
‘Satina’	3.33	0.83
‘Diana’	7.83	4.67
‘Theresa’	6.83	3.00

Potato chips quality was assessed by visually scoring the chips colour after deep frying of 1.2-2.0mm tuber slices in oil at 160-180°C for 2-3min (PUTZ, 1989), using a 1-9 colour scale. 1=very dark chips colour, very bad chips quality; 9=very light yellow chips colour, very good chips quality. CQA refers to the chips quality after harvest in autumn; CQS refers to the chips quality in spring after tuber cold storage for 3-4 months.

Out of 80 cloned cDNA sequences per genotype, a sequence was defined as an allele when it was found twice in two independent PCRs. Additionally, the consensus sequence of all alleles found in one genotype was used for allele definition when variable sequence polymorphisms occurred.

Pain-I invertase alleles obtained from each genotype are listed in Table 3.1.3.

Table 3.1.3: Overview of *Pain-I* alleles.

Genotype	Full-length clones	<i>Pain-I</i> alleles
‘Satina’	9	<i>Pain_SA</i>
		<i>Pain_SN</i>
‘Diana’	16	<i>Pain_DA</i>
		<i>Pain_DN1</i>
		<i>Pain_DN2</i>
‘Theresa’	8	<i>Pain_TN1</i>
		<i>Pain_TN2</i>
P18	7	<i>Pain_P18N1</i>
		<i>Pain_P18N2</i>
P40	8	<i>Pain_P40N1</i>
		<i>Pain_P40N2</i>
P54	6	<i>Pain_P54N</i>

The full-length clone number refers to the number of fully sequenced clones from each genotype used for allele definition. Allele names refer to the identification of associated *Pain-I* alleles, described in section 3.1.1.2. The ‘A’ in the allele name stands for ‘association with better potato chips quality’, and refers to clones containing SNP 1544. The ‘N’ in the allele name means ‘not associated’ with better potato chips quality.

From all six genotypes analyzed in this study, 12 *Pain-I* alleles were identified.

3.1.1.1.1 cDNA alleles of the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’¹

From nine full-length cDNA clones of the genotype ‘Satina’ two different alleles *Pain_SA* and *Pain_SN* were defined. Cloning and sequencing of ‘Diana’ cDNA resulted in 16 full-length clones, from which three different *Pain-I* alleles *Pain_DA*, *Pain_DN1*, and *Pain_DN2* were identified. For the cultivar ‘Theresa’ eight full-length clones were obtained. Two different alleles *Pain_TN1* and *Pain_TN2* were defined.

¹ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: ‘Satina’ (Appendix A 3.1.1), ‘Diana’ (Appendix A 3.1.2), ‘Theresa’ (Appendix A 3.1.3). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *Pain_SA* (Appendix A 3.1.4), *Pain_SN* (Appendix A 3.1.5), *Pain_DA* (Appendix A 3.1.6), *Pain_DN1* (Appendix A 3.1.7), *Pain_DN2* (Appendix A 3.1.8), *Pain_TN1* (Appendix A 3.1.9), *Pain_TN2* (Appendix A 3.1.10).

Comparing all seven alleles at the nucleotide level (Appendix A 3.1.11), 26 single nucleotide polymorphisms (SNPs) were detected. Ten of them resulted in an amino acid exchange (Table 3.1.4; Figure 3.1.1).

Table 3.1.4: SNPs present in ‘Satina’, ‘Diana’, and ‘Theresa’ alleles.

Position of cDNA SNP	<i>Pain_SA</i>	<i>Pain_SN</i>	<i>Pain_DA</i>	<i>Pain_DN1</i>	<i>Pain_DN2</i>	<i>Pain_TN1</i>	<i>Pain_TN2</i>	aa
32	C	T	C	T	C	T	C	C/T P11L
75	C	T	C	T	C	T	C	s.
174	G	A	G	A	G	A	G	s.
213	A	G	G	G	G	G	G	s.
280	G	G	A	G	G	G	G	A/G I94V
528	T	A	T	A	T	A	T	s.
552	C	T	C	T	T	T	T	s.
612	A	A	A	A	G	A	G	s.
718	A	G	A	G	G	G	G	A/G I240V
741	T	C	C	C	C	C	C	s.
777	T	T	T	T	C	T	C	s.
1068	C	C	C	C	G	C	G	C/G N356K
1212	G	G	G	G	G	G	A	s.
1316	C	C	T	C	C	C	C	T/C V439A
1143	C	G	G	G	G	G	G	s.
1544	A	C	A	C	C	C	C	A/C K515T
1574	A	A	A	A	T	A	T	A/T Y525F
1596	T	C	T	C	T	C	T	s.
1629	T	C	T	C	T	C	T	s.
1661	A	A	A	A	G	A	G	A/G Q554R
1689	G	G	A	G	G	G	G	s.
1749	T	T	C	T	T	T	T	s.
1830	C	C	C	C	C	C	T	s.
1843	G	T	G	T	G	T	G	T/G S615A
1857	C	C	T	C	C	C	C	s.
1895	G	A	G	A	G	A	G	A/G Q632R

SNP position numbering refers to cDNA sequence where ‘1’ represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are orange coloured. SNP 1544 (blue) was used to assign the associated SSCP fragments to a cloned allele (section 3.1.1.2) as well as for pyrosequencing analysis (section 3.1.2.1).

The two alleles of ‘Satina’ had a total of 14 SNPs, of which five caused an amino acid exchange. In the three ‘Diana’ allelic sequences 21 SNPs occurred, from which ten resulted in an amino acid substitution. The two ‘Theresa’ alleles differed in 15 SNPs. Six of them led to amino acid exchanges.

The amino acid alignment highlights the differences of all seven *Pain-1* alleles from the three tetraploid potato cultivars (Figure 3.1.1). The comparison of these deduced protein sequences revealed ten variable amino acid positions in the different genotypes.

Pain_DN1	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_TN1	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_SN	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_DN2	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_TN2	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_SA	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_DA	:	MATQYHSSYD	LE	NSASHY	TFLPDQ	PDSGHRKSLKII	ISGIFLSS	FLLLSVAFFP	ILNNQSPDLQ	SNRSRSPAPPS	RGVSSQGVSDK	:	83
Pain_DN1	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_TN1	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_SN	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_DN2	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_TN2	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_SA	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_DA	:	TFRD	VVNASH	VS	YAWS	NAMLS	WQRTAY	HFQPK	QNMND	PNGLY	HKGWYHL	LFYQYN	PD
Pain_DN1	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_TN1	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_SN	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_DN2	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_TN2	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_SA	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_DA	:	AMVP	DQWY	DING	VWTGS	ATILPD	GQIMMLY	TGTD	DDYV	QVQNL	AYPTNL	SDPL	LLD
Pain_DN1	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_TN1	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_SN	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_DN2	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_TN2	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_SA	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_DA	:	WTGP	QNGQ	WLLT	IGSK	IGKT	GIAL	VYETS	SNFTS	FKLL	DEV	LH	AV
Pain_DN1	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_TN1	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_SN	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_DN2	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_TN2	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_SA	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_DA	:	DDNK	QDHY	AIGT	YDLT	KNKW	TPD	PE	LD	CG	IG	LK	DY
Pain_DN1	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_TN1	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_SN	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_DN2	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_TN2	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_SA	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_DA	:	VLYD	KKTG	THLL	QWPVEE	IESLR	AG	DP	IV	KQ	VNL	QPGS	IELL
Pain_DN1	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_TN1	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_SN	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_DN2	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_TN2	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_SA	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_DA	:	SRG	ILG	PPF	GVV	VIADQ	TL	SEL	TPVY	FY	ISK	GAD	GRA
Pain_DN1	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_TN1	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_SN	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_DN2	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_TN2	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_SA	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K
Pain_DA	:	FAQ	GGR	T	V	T	S	R	I	Y	P	T	K

Figure 3.1.1: Amino acid alignment of *Pain-1* alleles from the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’. Amino acid exchanges are highlighted in colour.

Comparison of allelic cDNA sequences of the *Pain-1* gene from the three tetraploid cultivars revealed that different genotypes harbour amino acid sequence identical alleles. ‘Satina’, ‘Diana’, and ‘Theresa’ share one identical allele (*Pain_SN*=*Pain_DN1*=*Pain_TN1*). Furthermore, ‘Diana’, and ‘Theresa’ contain another allele identical at amino acid level (*Pain_DN2*=*Pain_TN2*).

3.1.1.1.2 cDNA alleles of the diploid potato genotypes P18, P40, and P54²

Cloning and sequencing of P18 cDNA resulted in seven full-length clones, from which two different alleles *Pain_P18N1* and *Pain_P18N2* were identified. In the case of P40 two alleles *Pain_P40N1* and *Pain_P40N2* out of eight full-length clones were found. From the genotype P54 six full-length clones were obtained. All six clones exhibited the same amino acid sequence. The P54 allele was named *Pain_P54N*.

The nucleotide sequence comparison (Appendix A 3.1.20) of the alleles described above detected 47 SNPs, from which 20 resulted in amino acid exchanges (Table 3.1.5; Figure 3.1.2).

Table 3.1.5: SNPs present in P18, P40, and P54 alleles.

Position of cDNA SNP	<i>Pain_P18N1</i>	<i>Pain_P18N2</i>	<i>Pain_P40N1</i>	<i>Pain_P40N2</i>	<i>Pain_P54N</i>	aa
18	T	T	C	C	T	s.
21	C	C	A	A	C	s.
32	C	T	C	C	T	C/T P11L
74	C	C	A	A	C	C/A P25H
75	C	C	C	C	T	s.
93	G	G	A	A	G	s.
97	C	C	A	C	C	A/C I33L
130	T	T	C	C	T	T/C F44L
145	G	G	T	T	G	G/T V49L
149	C	C	T	T	C	C/T A50V
174	G	A	G	G	A	s.
189	G	G	A	A	G	s.
196	T	T	G	G	T	T/G S66A
200	G	G	A	G	G	A/G H67R

² Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: P18 (Appendix A 3.1.12), P40 (Appendix A 3.1.13), P54 (Appendix A 3.1.14). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *Pain_P18N1* (Appendix A 3.1.15), *Pain_P18N2* (Appendix A 3.1.16), *Pain_P40N1* (Appendix A 3.1.17), *Pain_P40N2* (Appendix A 3.1.18), *Pain_P54N* (Appendix A 3.1.19).

Position of cDNA SNP	<i>Pain_P18N1</i>	<i>Pain_P18N2</i>	<i>Pain_P40N1</i>	<i>Pain_P40N2</i>	<i>Pain_P54N</i>	aa
280	A	G	G	G	G	A/G I94V
369	T	T	T	C	T	s.
528	T	A	T	T	A	s.
534	T	T	T	G	T	s.
552	C	T	T	T	T	s.
591	C	C	T	T	C	s.
612	A	A	G	G	A	s.
637	A	A	G	G	A	T/A T213A
718	A	G	G	G	G	A/G I240V
723	G	A	G	G	G	s.
834	C	C	C	A	C	s.
852	G	G	A	G	G	s.
927	T	T	T	C	T	s.
1050	C	T	C	C	C	s.
1161	A	A	G	A	A	s.
1267	G	G	A	G	G	A/G R423G
1316	T	C	C	C	C	T/C V439A
1340	T	T	C	C	T	T/C V447A
1359	A	A	A	G	A	s.
1522	G	G	G	A	G	G/A V508I
1544	A	C	C	C	C	A/C K515T
1596	T	C	T	T	C	s.
1603	G	G	C	G	G	C/G Q535E
1629	T	C	T	T	C	s.
1640	A	A	T	T	A	A/T E547V
1683	C	C	C	G	C	s.
1689	A	G	G	G	G	s.
1698	G	A	G	G	A	s.
1749	C	T	T	T	T	s.
1776	G	G	A	G	G	s.
1843	G	T	G	G	T	G/T A615S
1857	T	C	C	C	C	s.
1895	G	A	G	G	A	G/A R632Q

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are orange coloured. SNP 1544 (blue) was used to assign the associated SSCP fragments to a cloned allele (section 3.1.1.2) as well as for pyrosequencing analysis (section 3.1.2.1).

From 17 SNPs of the two P18 alleles, seven caused an amino acid exchange. In the two P40 allelic sequences 14 SNPs occurred, from which five resulted in an amino acid substitution.

		*	420	*	440	*	460	*	480										
Pain_DA	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_P18N1	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_SA	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_DN2	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_TN2	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_SN	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_P54N	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_P18N2	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_TN1	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_DN1	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_P40N1	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486
Pain_P40N2	:	WASVQSI	PRTVL	YDKKT	GTHLLQWP	VVEEIES	LRAGDP	IVKQV	NLQPGS	IELLHV	DSAAEL	DI	EASF	FEVDK	VALQ	GI	EADH	:	486

		*	500	*	520	*	540	*	560	
Pain_DA	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_P18N1	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_SA	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_DN2	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_TN2	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_SN	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_P54N	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_P18N2	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_TN1	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_DN1	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_P40N1	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567
Pain_P40N2	:	VGFSCSTSGGAASRGILGPF	VVVIADQ	KLSELT	TPVYFY	ISKGADGRAE	THFCADQTRSS	EAPGVAK	QVYSSVPVLDGEK	: 567

		*	580	*	600	*	620	*	
Pain_DA	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_P18N1	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_SA	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_DN2	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_TN2	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_SN	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_P54N	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_P18N2	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_TN1	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_DN1	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_P40N1	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			
Pain_P40N2	:	HSMRLLVDHSIVESFAQGGRTVITSRIYPTKAVNGAARLFVFN	NATGASVTASVKIWSLESANIR	SFPLQDL	:	639			

Figure 3.1.3: Amino acid alignment of all cloned *Pain-1* invertase alleles. Amino acid exchanges are highlighted in colour.

Protein sequence comparison of the deduced *Pain-1* alleles from all analyzed genotypes showed that amino acids differed at 23 positions, of which five were genotype specific and occurred only once.

3.1.1.1.4 Phenetic trees of all *Pain-1* invertase alleles of the analyzed potato genotypes

In addition to the multiple amino acid alignment (section 3.1.1.1.3), the phenetic tree analysis was applied to group the invertase alleles according to similarity at amino acid as well as at nucleotide level. Using the neighbour-joining method, the allelic classification visualized that *Pain-1* alleles from all analyzed potato genotypes grouped in two clades and four to five subclades (Figure 3.1.4, Figure 3.1.5).

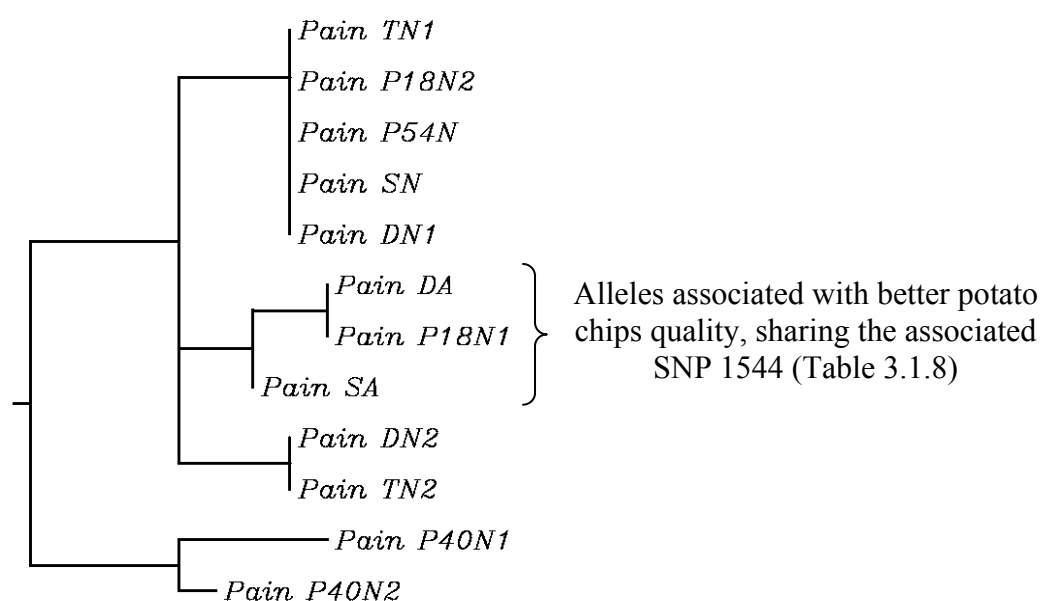


Figure 3.1.4: Amino acid based phenetic tree (Neighbour-joining method) of all cloned *Pain-1* invertase alleles.

The first clade includes alleles from the genotypes ‘Satina’, ‘Diana’, ‘Theresa’, P18, and P54. The first subclade of the first clade consists of five alleles from five different genotypes. The alleles *Pain_SN*, *Pain_DN1*, *Pain_TN1*, *Pain_P18N2*, and *Pain_P54N* have the same amino acid sequence, although polymorphisms in the corresponding nucleotide sequences occurred (Figure 3.1.5).

The second subclade consists of the alleles *Pain_SA* and *Pain_DA*, which are associated with better potato chips quality. The two alleles are not identical at the amino acid level but possess cDNA SNP 1544 that is associated with better potato chips quality (section 3.1.1.2). The allele *Pain_P18N1* of the diploid genotype P18 has an identical amino acid sequence to *Pain_DA*.

The third subclade includes the two alleles *Pain_DN2* and *Pain_TN2*, which have the same amino acid sequence.

The second main clade consists of the two outlying P40 alleles *Pain_P40N1* and *Pain_P40N2*. The two alleles show the highest diversity compared to the other cloned alleles.

Cloning and sequencing of *Pain-1* alleles showed that tetraploid and diploid genotypes contain alleles identical in their amino acid sequence but different at nucleotide level. The following Table 3.1.6 summarizes the nucleotide comparison of amino acid identical alleles of different genotypes. The allelic nucleotide sequence was defined based on the consensus sequence of multiple alignment of full-length clones obtained from each genotype (Table 3.1.3). Although SNPs are present at positions in the corresponding cDNA, the nucleotide polymorphisms resulted in one and the same amino acid sequence.

Table 3.1.6: Genotype specific nucleotide differences of alleles identical at amino acid level (Appendix A 3.1.21).

A) Comparison of the alleles *Pain_SN*, *Pain_DN1*, *Pain_TN1*, *Pain_P18N2*, and *Pain_P54N*.

Position of cDNA SNP	<i>Pain_SN</i>	<i>Pain_DN1</i>	<i>Pain_TN1</i>	<i>Pain_P18N2</i>	<i>Pain_P54N</i>
75	T	T	T	C	T
486	A	G	G	G	G
723	G	G	G	A	G
1050	C	C	C	T	C
1236	T	C	T	T	T

B) Alleles of the alleles *Pain_DA* and *Pain_P18N1*.

Position of cDNA SNP	<i>Pain_DA</i>	<i>Pain_P18N1</i>
816	G	A
1698	A	G

C) Comparison of the alleles *Pain_DN2* and *Pain_TN2*.

Position of cDNA SNP	<i>Pain_DN2</i>	<i>Pain_TN2</i>
1830	C	T

The nucleotide polymorphisms between all *Pain-1* (Appendix A 3.1.21) were visualized using the phenetic tree analysis (Figure 3.1.5).

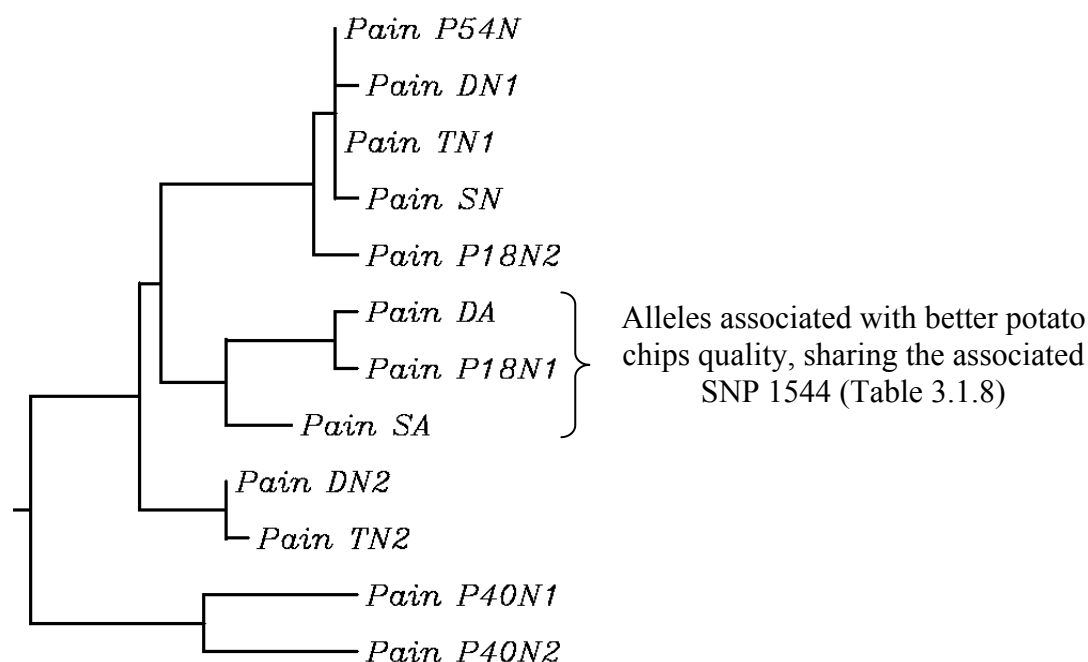


Figure 3.1.5: Nucleotide sequence based phenetic tree (Neighbour-joining method) of all cloned *Pain-1* invertase alleles.

Allele specific SNPs were detected in at least two sequences and used for comparing the allelic nucleotide polymorphisms. The two phenetic trees are very similar, just displaying more subclades due to a higher number of nucleotide polymorphisms compared to the amino acid exchanges (Figure 3.1.4).

3.1.1.2 Identification of associated *Pain-1* alleles

The *Pain-1* locus maps to potato chromosome III (CHEN ET AL., 2001) in a region associated with tuber quality traits where a QTL for potato tuber sugar content, *Sug3a*, was identified (MENÉNDEZ ET AL., 2002). In the latter study, the *Pain-1* gene was not mapped directly in the mapping population used, but was considered as a candidate gene for the QTL *Sug3a*.

Single-strand conformation polymorphism (SSCP) analysis revealed an association of *Pain-1* SSCP fragments with starch and sugar content of potato tubers. SSCP fragments found to be associated with better potato chips quality were named *Pain1-5c*, *Pain1-8c* and *Pain1-9a* (LI ET AL., 2008). The occurrence of these fragments in the genotypes was highly correlated indicating that the three fragments are in nearly absolute LD, and suggesting that these allelic fragments correspond to the same associated *Pain-1* invertase allele. The associated SSCP fragments are present in the cultivars ‘Satina’, ‘Diana’, and absent in ‘Theresa’ (Table 3.1.7).

Table 3.1.7: Associated SSCP fragments *Pain1-5c*, *Pain1-8c* and *Pain1-9a* present in ‘Satina’, ‘Diana’, and ‘Theresa’.

Genotype	<i>Pain1-5c</i>	<i>Pain1-8c</i>	<i>Pain1-9a</i>
‘Satina’	0	1	1
‘Diana’	1	1	1
‘Theresa’	0	0	0

0=SSCP fragment is absent, 1=SSCP fragment is present.

To assign a cloned invertase allele (section 3.1.1.1) to the corresponding associated SSCP fragments, two groups, each consisting of 15 potato genotypes that were scored either positive (present) or negative (absent) for the associated SSCP fragments, were used to identify the corresponding SNPs.

Based on multiple nucleotide alignment (Appendix A 3.1.22) of the three cultivars ‘Satina’, ‘Diana’, and ‘Theresa’, the cDNA SNP 1544 was selected present in ‘Satina’ and ‘Diana’, and absent in ‘Theresa’ referring to the SSCP fragment distribution (Table 3.1.7). The distribution of SNP 1544, which causes an amino acid exchange of threonine (T) in the non associated to lysine (K) in the associated alleles is shown in Table 3.1.8.

Table 3.1.8: Comparison and distribution of associated SSCP fragments and SNP 1544.

A) Genotypes scored negative for associated SSCP fragments

Genotype	SSCP			SNP 1544
	<i>Pain1-5c</i>	<i>Pain1-8c</i>	<i>Pain1-9a</i>	
‘Leyla’ (St01)	0	0	0	C
‘Marabel’ (St02)	0	0	0	C
‘Solara’ (St03)	0	0	0	C
‘Vitará’ (St04)	0	0	0	C
‘Ponto’ (St06)	0	0	0	C
‘Marlen’ (St08)	0	0	0	C
‘Eldena’ (St09)	0	0	0	C
‘Theresa’ (St10)	0	0	0	C
‘Goldika’ (St11)	0	0	0	C
‘Saturna’ (St12)	0	0	0	C
‘Karlén’ (St14)	0	2	0	C
‘Kolibri’ (St15)	0	2	0	C
‘Terra’ (St18)	0	0	0	C
‘Solist’ (St19)	0	2	0	C
‘Melina’ (St20)	0	2	0	C

B) Genotypes scored positive for associated SSCP fragments

Genotype	SSCP			SNP 1544
	<i>Pain1-5c</i>	<i>Pain1-8c</i>	<i>Pain1-9a</i>	
‘Milva’ (St05)	1	0	1	C/A
‘Tomensa’ (St07)	1	0	1	C/A
‘Fasan’ (St16)	1	2	1	C
‘Apart’ (St24)	0	1	1	C/A
‘Diana’ (St28)	1	1	1	C/A
‘Orlando’ (St31)	1	0	1	C/A
‘Satina’ (St33)	0	1	1	C/A
B05	1	1	1	C/A
B07	1	0	1	C/A
B16	1	1	1	C
B17	1	1	1	C/A
B30	1	0	1	C
B32	1	1	1	C/A
B38	1	1	1	C/A

St=potato cultivars defined as standards, B=BNA breeding clones, 0=SSCP fragment is absent, 1=SSCP fragment is present, 2=SSCP fragment could not be analyzed properly, SNP 1544=analyzed SNP for association with better potato chips quality, C=cytosine, A=adenine.

All genotypes, which scored negative for the associated SSCP fragments (Table 3.1.8 A), contain the nucleotide C at cDNA SNP position 1544. Genotypes, which scored positive for associated SSCP fragments (Table 3.1.8 B), show a C/A-polymorphism nearly equally distributed as the corresponding SSCP fragments. Three out of 14 analyzed genotypes did not display the C/A-polymorphism.

For further analysis and molecular description of the cloned *Pain-1* alleles, the term ‘associated with better potato chips quality’ is used for alleles of phenotypically characterized genotypes displaying the C/A-polymorphism at cDNA position 1544.

This polymorphism results in an amino acid substitution at position 515 of the corresponding protein sequence, exchanging threonine (T) in the non associated invertase alleles to lysine (K) in the associated *Pain-1* alleles. To study the effects of the amino acid differences of cloned *Pain-1* alleles, analysis was continued with 3D-modelling (section 3.1.1.3) and functional characterization (section 3.1.2).

3.1.1.3 Three-dimensional modelling of *Pain-1* alleles

3D-modelling was performed by Pawel Durek, MPIMP/Golm. The modelling of the allelic invertase molecular structure was based on the invertase 3D and crystal structure of cyanobacteria (ALBERTO ET AL., 2004). The models are comparative, superimposing two allelic sequences. Differences of the alleles are highlighted. The modelling was applied for associated and non associated *Pain-1* alleles of the tetraploid potato cultivars ‘Satina’ and ‘Diana’. The models include the putative sucrose binding site with the substrate sucrose. In

addition to the structural visualization of amino acid exchanges, also the electrostatic potential (EP) of the molecules was mapped at pH 4.7 mimicking the vacuolar conditions.

3.1.1.3.1 Structural modelling of alleles of the cultivars ‘Satina’ and ‘Diana’

In the first analysis of allelic molecular structures, the two ‘Satina’ alleles *Pain_SA* and *Pain_SN*, and the three ‘Diana’ alleles *Pain_DA*, *Pain_DN1*, and *Pain_DN2* were used. The models compare the allelic sequences with each other, meaning that one sequence superimposes the other and vice versa. Based on multiple alignment (Figure 3.1.1) of the allelic protein sequences, regions, which are affected directly or indirectly by amino acid exchanges, were identified (Figure 3.1.6; Figure 3.1.7).

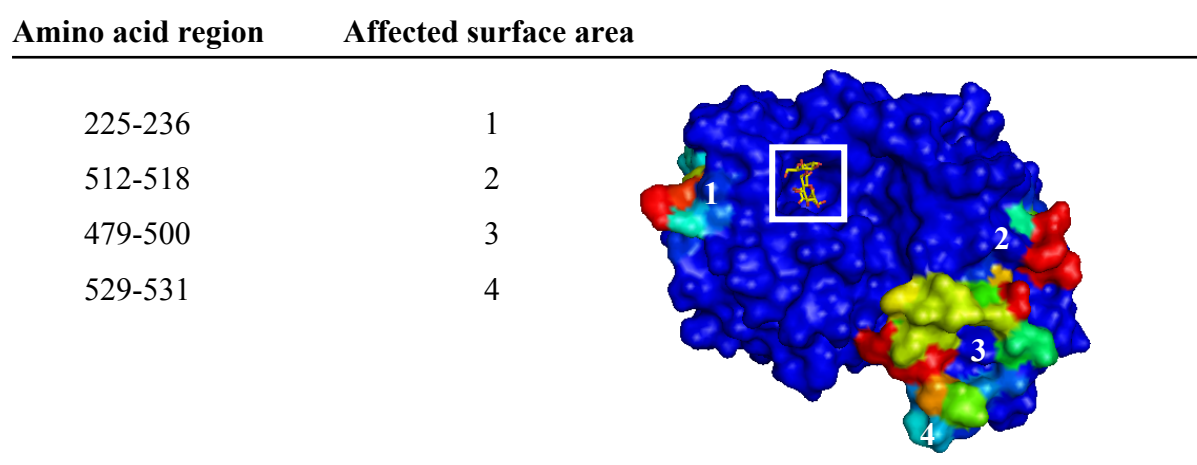


Figure 3.1.6: Correlation of amino acid regions or exchanges and affected surface areas of *Pain-1* alleles. Amino acids are numbered based on the multiple sequence alignment (Figure 3.1.1). Areas are numbered from 1-4. Red: strong structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from red (strong) to blue (weak). The putative sucrose binding site with the substrate sucrose is framed.

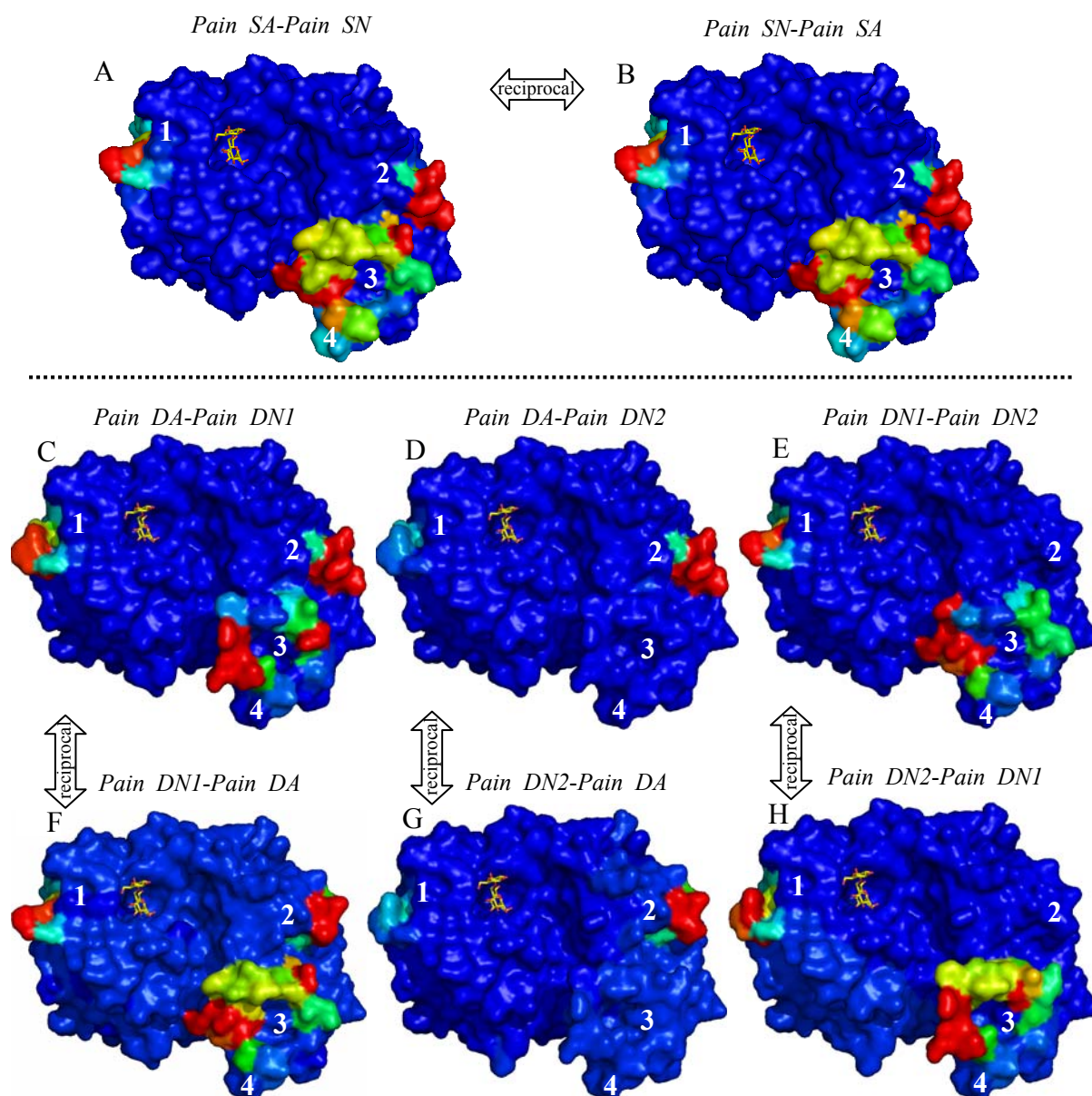


Figure 3.1.7: Structural comparison of ‘Satina’ alleles *Pain_SA*, *Pain_SN*, and ‘Diana’ alleles *Pain_DA*, *Pain_DN1*, *Pain_DN2*. **A:** *Pain_SA* superimposes *Pain_SN*. **B:** *Pain_SN* superimposes *Pain_SA*. **C:** *Pain_DA* superimposes *Pain_DN1*; **D:** *Pain_DA* superimposes *Pain_DN2*; **E:** *Pain_DN1* superimposes *Pain_DN2*; **F:** *Pain_DN1* superimposes *Pain_DA*; **G:** *Pain_DN2* superimposes *Pain_DA*; **H:** *Pain_DN2* superimposes *Pain_DN1*. Red: strong structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from red (strong) to blue (weak).

The models of the invertase molecules of the alleles *Pain_SA* and *Pain_SN* and *Pain_DA*, *Pain_DN1*, and *Pain_DN2* showed structural differences on the enzyme’s surface. All analyzed molecules possess similar affected surface areas (area definition: Figure 3.1.6). Surface area 1 differs in all represented models, but is less distinctive between the alleles *Pain_DA* and *Pain_DN2* (Figure 3.1.7 D, G). Region 1, defined by the amino acids 225-236, manifests sterical differences although amino acids in the corresponding region do not differ between the alleles. The same observations were made for the surface areas 3 and 4, defined by amino acids 479-500 and 529-531 respectively, where the alleles display identical amino

acid sequences. These areas are affected indirectly by other amino acid substitutions leading to a variable folding of the protein. Accountable amino acids are subject of ongoing investigations.

Region 2 is defined by amino acids 512-518 showing a lysine (K) in the associated alleles *Pain_SA* and *Pain_DA*, and a threonine (T) in the alleles *Pain_SN*, *Pain_DN1*, and *Pain_DN2* at position 515. The corresponding nucleotide SNP 1544 was found to be associated with better potato chips quality (section 3.1.1.2). The structural modification 2 is absent in the models E and H (Figure 3.1.7) because of the absence of SNP 1544 in the superimposed alleles *Pain_DN1* and *Pain_DN2*.

The structural effects caused by allelic sequences resulted in invertase folding variations of the *Pain-I* molecules. The function of all identified variable surface areas is unknown.

3.1.1.3.2 Modelling the electrostatic potential (EP) of alleles of the cultivars ‘Satina’ and ‘Diana’

The mapping of the EP of the *Pain-I* alleles *Pain_SA*, *Pain_SN*, *Pain_DA*, *Pain_DN1*, and *Pain_DN2* revealed charge differences of the molecules at pH 4.7 (Figure 3.1.8).

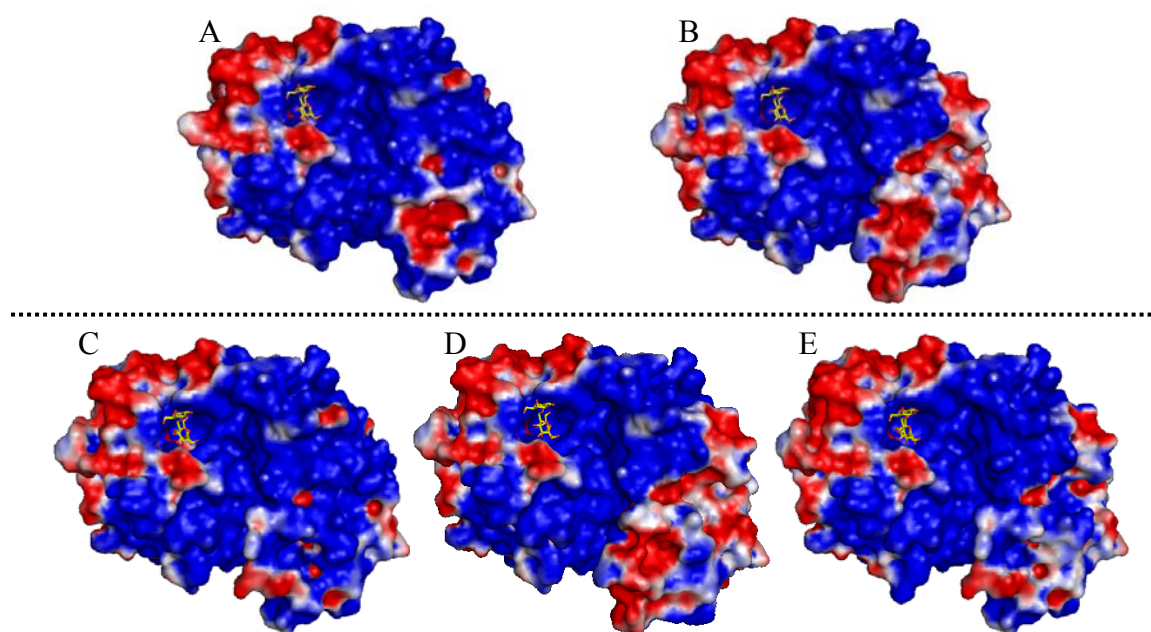


Figure 3.1.8: EP of the alleles *Pain_SA*, *Pain_SN*, *Pain_DA*, *Pain_DN1*, and *Pain_DN2*. A: *Pain_SA*. B: *Pain_SN*. C: *Pain_DA*; D: *Pain_DN1*; E: *Pain_DN2*. Red: negatively charged; blue: positively charged; white: neutrally charged.

The EP modelling of the two alleles from the cultivar ‘Satina’ showed that *Pain_SA* is more positively charged than *Pain_SN* (Figure 3.1.8 A, B). The three ‘Diana’ alleles also exhibit charge differences to each other. The allele *Pain_DN1* (Figure 3.1.8 D) possesses an extended

negative charge in comparison to the other two alleles (C, E). The EP of the allele *Pain_DA* (C) showed more positive charges in regions where allele *Pain_DN2* was more neutral.

Focusing on the EP of the putative sucrose binding site

The putative sucrose binding site (Figure 3.1.6) is positively charged matching the partial negative charge of the substrate sucrose due to the hydroxyl groups.

Zooming into the putative sucrose binding domain revealed charge differences between the molecules *Pain_SA* (A), *Pain_SN* (B), *Pain_DA* (C), *Pain_DN1* (D), and *Pain_DN2* (E); (Figure 3.1.9).

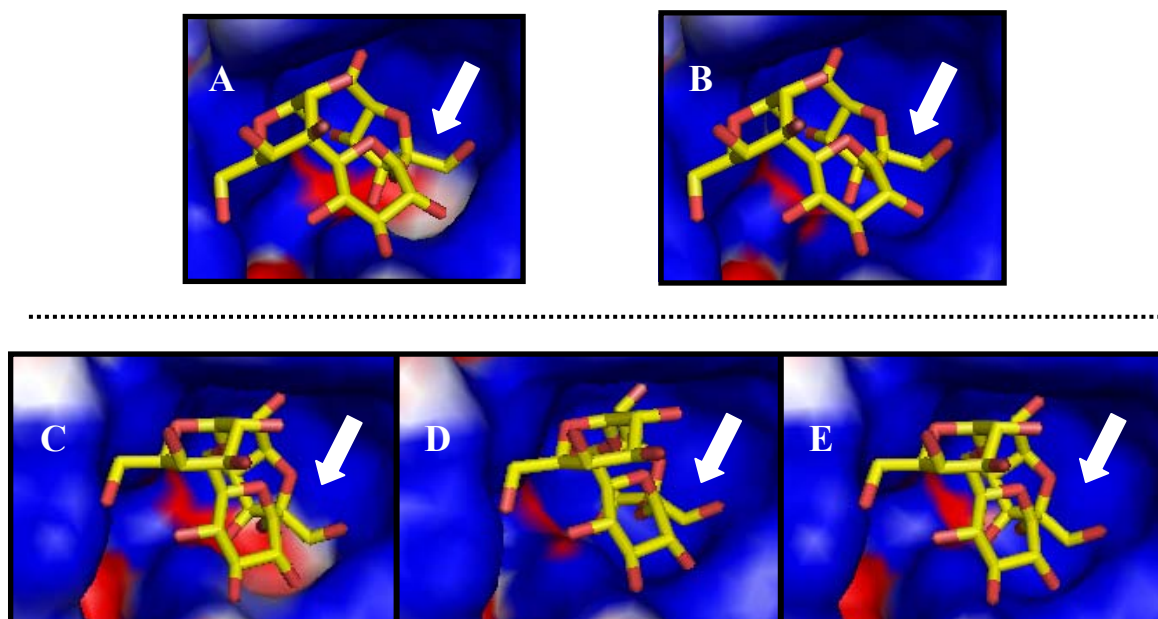


Figure 3.1.9: Focusing of the EP of the putative sucrose binding site of the alleles of the cultivars ‘Satina’ and ‘Diana’. A: *Pain_SA*; B: *Pain_SN*; C: *Pain_DA*, D: *Pain_DN1*, E: *Pain_DN2*. Red: negatively charged; blue: positively charged; white: neutrally charged.

Close-up views of the putative sucrose binding domain revealed strong charge changes between the molecules *Pain_SA* (A) and *Pain_SN* (B) (Figure 3.1.9). The site of the associated allele *Pain_SA* is characterized by a neutral to negative charge composition compared to the positive charge of the allele *Pain_SN*. The EP of the putative sucrose binding site of the three ‘Diana’ alleles also showed apparent changes of charge. The EP of the associated allele *Pain_DA* (C) switched from a positive to a neutral/negative charge. The charge of the alleles *Pain_DN1* (D), and *Pain_DN2* (E) differed not visibly.

3.1.1.3.3 Overview of 3D-modelling analysis of *Pain-1* alleles

3D-modelling is a tool to gain first insights in the possible consequences of different invertase alleles. The models nicely show that amino acid exchanges manifest on the surface of the enzyme. However, no effect on the structure of the putative sucrose binding domain was

detected in any of the modelled molecules. The characterization of the EP showed differences in charge among the analyzed proteins. A dramatic charge difference of the putative sucrose binding site was observed in the associated alleles *Pain_SA* and *Pain_DA* (Figure 3.1.9 A, C). The EP switched from positive to neutral. The EP changes could not be correlated to amino acid exchanges nearby the sucrose binding site. The causative amino acids were not yet analyzed, but are subject of ongoing investigations in the research project.

3.1.1.4 Genomic organization of the *Pain-1* locus

❖ BAC library screens

High density BAC library screens with two different PCR generated probes were performed to determine the genomic sequence of the *Pain-1* locus. Using the primers Pain1-5f/Pain1-5r (LI ET AL., 2005; chapter 2, Table 2.1.10A), Probe 1 was generated, which consists of exon and intron based sequences of *Pain-1*. Probe 2 only consists of exon based sequences and was amplified with the primers Pain_SondF1/Pain_SondR (chapter 2, Table 2.1.10A).

The screening of two different BAC libraries (BA and BC, BALLVORA ET AL., 2002, 2007) constructed with genomic DNA of the same diploid genotype resulted in 21 positive BAC clones (Table 3.1.9).

Table 3.1.9: Positive BAC clones.

Library	BAC clones	
BA BAC library	BA1: Plate 32N19	
	BA2: Plate 128A5	
	BA3: Plate 213B14	
BC BAC library	BC1: Plate 14O7	BC10: Plate 47C9
	BC2: Plate 21K3	BC11: Plate 57A16
	BC3: Plate 27G10	BC12: Plate 57P6
	BC4: Plate 32J7	BC13: Plate 68K22
	BC5: Plate 34G11	BC14: Plate 149O15
	BC6: Plate 34O24	BC15: Plate 216L1
	BC7: Plate 35N9	BC16: Plate 223N21
	BC8: Plate 35E21	BC17: Plate 239N10
	BC9: Plate 45P17	BC18: Plate 244C12

BACs are numbered and their position in the *E. coli* microtiter plates is listed.

Following the hybridisation, clones from the BC library for sequencing were selected based on the results of PCR, colony-lift, and Southern blot. Out of 18 hybridization positive *Pain-1* BC BAC clones, eight clones were found positive for full-length PCR amplification using

Pain-1 specific primers. All analyzed clones were colony-lift positive and eight BACs were Southern blot positive (Table 3.1.10; Figure 3.1.10).

Table 3.1.10: Summary of the results for BAC insert sequencing.

BC clones	Filter hybridisation	PCR	Colony-lift	Southern blot
BC1: Plate 14O7	yes	no	yes	no
BC2: Plate 21K3	yes	no	yes	yes
BC3: Plate 27G10	yes	no	yes	yes
BC4: Plate 32J7	yes	no	yes	no
BC5: Plate 34G11	yes	yes	yes	yes
BC6: Plate 34O24	yes	no	yes	no
BC7: Plate 35N9	yes	no	yes	no
BC8: Plate 35E21	yes	no	yes	yes
BC9: Plate 45P17	yes	yes	not analyzed	not analyzed
BC10: Plate 47C9	yes	no	yes	no
BC11: Plate 57A16	yes	no	yes	not analyzed
BC12: Plate 57P6	yes	no	yes	no
BC13: Plate 68K22	yes	no	yes	no
BC14: Plate 149O15	yes	no	yes	no
BC15: Plate 216L1	yes	no	yes	no
BC16: Plate 223N21	yes	yes	yes	yes
BC17: Plate 239N10	yes	no	yes	yes
BC18: Plate 244C12	yes	yes	yes	yes

For PCR reaction Probe 1 (Pain1-5f/Pain1-5r, chapter 2, Table 2.1.10A), Probe 2 (Pain_SondF1/Pain_SondR, chapter 2, Table 2.1.10A), or *Pain-1* full-length primers (ZrPain-F/PainUni-R, chapter 2, Table 2.1.2) were used. Colony-lift and Southern blot hybridisation were performed using Probe 1.

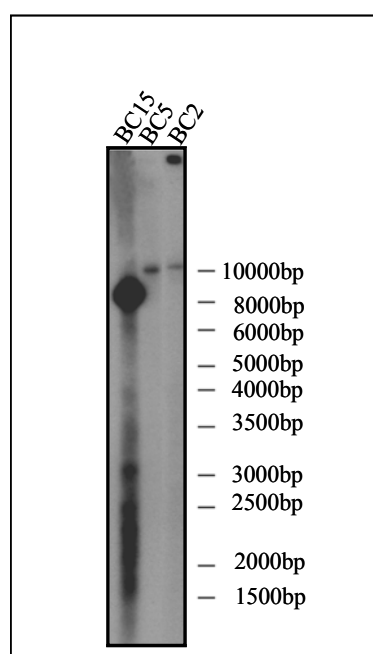


Figure 3.1.10: Southern blot analysis of *Sma*I digested BC BAC clones. The pattern of this first line was only represented by the BAC BC15. The restriction pattern of BC5 was also observed for the clones BC12 and BC14. The BC2 pattern was also detected for the BACs BC8 and BC13.

The *Pain-1* locus of BC library clones BC14 and BC17 was custom sequenced by GATC Biotech, Konstanz using primer walking as sequencing method. Full-length BAC insert sequencing was performed for the clones BC2 (insert size: 75kb), BC14 (insert size: 75kb), and BC15 (insert size: 97kb), which showed differential restriction patterns in Southern blot

analysis (Figure 3.1.10) suggesting different *Pain-I* alleles or gene loci. Full-length BAC insert sequencing was carried out by MWG Biotech AG, Ebersberg using the 454 sequencing technique on the GS FLX system. The BAC clones BC14 and BC15 as well as BC17 harboured the same genomic invertase allele. BAC clone BC2 was found invertase negative, meaning that no invertase sequence was detected.

The genomic sequences of the *Pain-I* gene received by different sequencing technologies differed in one to six nucleotides in the second intron of the gene. The sequences resulted from primer walking showed more nucleotide differences than sequences obtained by 454 sequencing.

❖ BAC Annotation

Full-length sequencing of the BAC inserts revealed sequence and structural information of *Pain-I* and of genes flanking the *Pain-I* locus. Since the BAC clones BC14 and BC15 contained the same genomic *Pain-I* invertase sequence, only the annotation of BC14 is shown (Table 3.1.11).

Table 3.1.11: BC14 sequence annotation.

Strand	Apollo name	Position on the BAC insert (bp)	Description
+	gene 10	9,298-13,989	putative retrotransposon protein
+	gene 20	15,377-16,656	unknown
+	gene 30	31,121-33,084	putative transposon protein
+	gene 40	33,625-35,078	putative transposase
+	gene 50	74,626-78,901	putative retroelement polyprotein
-	gene 60	6,138-5,488	putative reverse transcriptase
-	gene 70	61,166-57,216	invertase <i>Pain-I</i> , beta-fructofuranosidase (glucoside hydrolase family 32)

BAC annotation was carried out using the software Apollo Genome Annotation and Curation Tool, version 1.9.8. The invertase gene was named 'gene 70' in the Apollo BAC sequence characterization. The invertase gene is written in bold.

The screened BAC libraries BA and BC harbour genomic DNA of the same diploid genotype P6/210, which is a hybrid derived from the cross of the parental genotypes P40 x P41 (LEISTER ET AL., 1996). The genotype P40 was also selected in this study for invertase allele characterization. The sequence alignment of the two P40 cDNA alleles with the protein sequence of the gene 70 (*Pain-I*) from BAC BC14, which is identical to the *Pain-I* genomic sequences of the BACs BC 15 and BC17, showed no sequence identity (Appendix A 3.1.23). Therefore, the detected BAC allele for the gene *Pain-I* originates from the other parental genotype P41.

❖ Genomic structure of the *Pain-1* gene

The exon and intron organization of the *Pain-1* gene was determined using multiple sequence alignment of *Pain-1* cDNA alleles and the genomic *Pain-1* sequence of BAC BC14 (Appendix A 3.1.24³).

The *Pain-1* gene consists of seven exons and six introns and has a length of 3951bp (Figure 3.1.11).

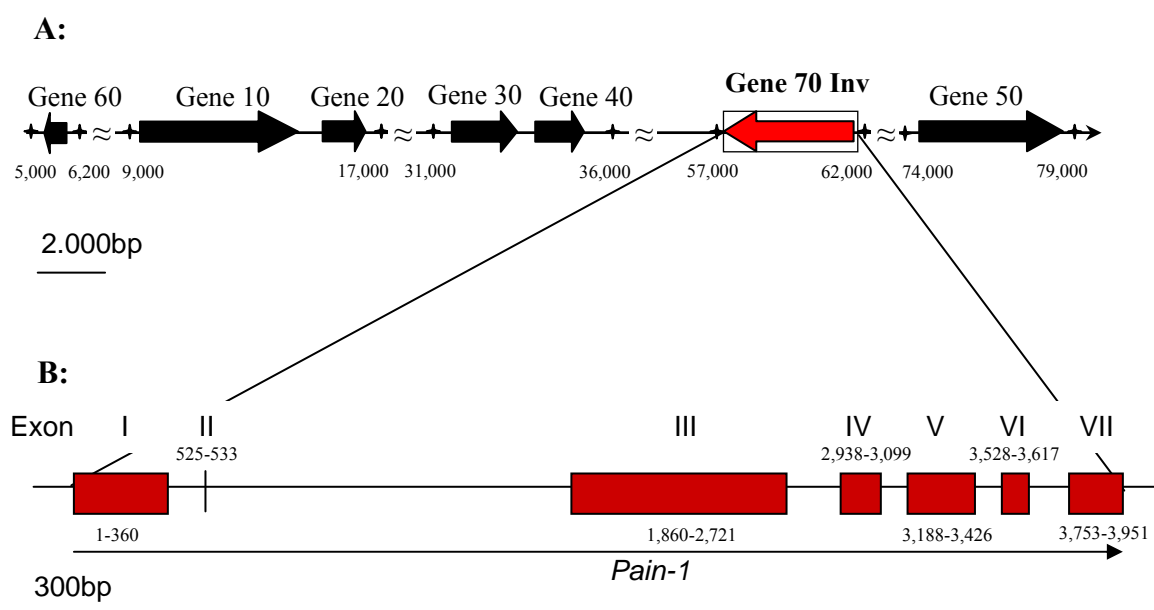


Figure 3.1.11: Position of the *Pain-1* gene within the sequenced BAC insert of BC14 (A) and genomic organization of the *Pain-1* gene (B). A: The direction of the annotated genes (Table 3.1.11) is indicated by arrows. \blacktriangleright : 5'-3' orientation (+ strand); \blacktriangleleft : 3'-5' orientation (- strand). The lengths are given in base pairs. B: Exons are drawn in red and numbered from I to VII, the length is given in base pairs. The arrow demonstrates the length of the gene without promoter and terminator sequences.

It is known from potato invertase loci encoding insoluble acid isoforms that two genes are linked in direct tandem repeat (MADDISON ET AL., 1999). The genes organized in this way are either separated by 2,3kb (MADDISON ET AL., 1999) or by 8kb (chapter 3.3, Figure 3.3.12). The insert of BAC BC14, which has a length of 75kb contained no second invertase gene. Within the BAC insert *Pain-1* gene surrounding sequences should allow to detect a probable tandem repeat organization if present (Figure 3.1.10 A). This possibility was excluded for the *Pain-1* locus.

The range of *Pain-1* exons and introns are summarized in Table 3.1.12.

³ The alignment program Multalin version 5.4.1 (<http://bioinfo.genetoul.fr>) used revealed problems in the comparison of the genomic *Pain-1* sequence with the *Pain-1* cDNA sequence. The mini-exon II was not aligned properly, one additional nucleotide occurred.

Table 3.1.12: Ranges of the exons and introns of the *Pain-1* gene.

Exon number	Range (bp)	Intron number	Range (bp)
I	1-360	I	361-524
II	525-533	II	534-1859
III	1,860-2,721	III	2,722-2,937
IV	2,938-3,099	IV	3,100-3,187
V	3,188-3,426	V	3,427-3,527
VI	3,528-3,617	VI	3,618-3,752
VII	3,753-3,951		

Plant invertases isolated to date have fairly similar structures and consist of six to eight exons (reviewed in TYMOWSKA-LALANNE & KREIS, 1998). The genes contain one extremely small exon (exon II), which only codes for the core tripeptide DPN of the conserved β -fructosidase motif NDPNG (TYMOWSKA-LALANNE & KREIS, 1998).

3.1.2 Functional characterization of the *Pain-1* gene

3.1.2.1 Differential expression analysis of *Pain-1* alleles during cold storage of potato tubers

Plant invertases are influenced by a variety of intra- and extracellular factors. It has been shown that invertases are regulated by temperature. ROREM & SCHWIMMER (1963) first demonstrated that temperature affects the level of invertase activity. Invertase activity is always detectable in potato tubers, but the highest level is measured when tubers have been stored at low temperature (PRESSEY, 1966). ZHOU ET AL. (1994) detected soluble acid invertase transcripts in potato tubers stored at 1°C, but not in those stored at 10°C. Strongly induced transcript accumulation of soluble acid invertase was also shown for tuber samples stored at 4°C after 7-10 days (ZRENNER ET AL., 1996; BAGNARESI ET AL., 2008).

Expression analysis was performed with two biological replicates of potato tubers stored for 1, 2, 3, and 4 weeks at 4°C in the dark as well as with potato tuber samples frozen in liquid nitrogen directly after harvest and kept at -80°C.

Two different strategies were used to define the allelic expression pattern. The first strategy was based on pyrosequencing analysis to separate the different *Pain-1* alleles within a genotype. The pyrosequencing analysis was performed using cDNA from tubers stored for 0-4 weeks at 4°C in the dark, and tuber genomic DNA to determine the allele dosage. Comparison of both samples revealed specific expression patterns not just for each genotype but also between different alleles of the same genotype. In the following Figures the relationship between the presence of alleles in the genome and their transcription levels are illustrated. Accordingly, both values are shown together in terms of the relative expression level.

Additionally, plasmids harbouring one allele of complementary SNPs of corresponding genotypes were mixed in different ratios to monitor the accuracy of pyrosequencing analysis. Plasmid based measurements, working as positive controls, showed that SNP depending variations of $\pm 5\%$ occurred. Values of cDNA and genomic dosages of the analysed alleles as determined by pyrosequencing were corrected for the observed SNP specific variations.

qRT-PCR was carried out as second strategy to measure the total amount of invertase transcript levels during the applied cold storage period of four weeks.

3.1.2.2.1 Expression pattern of *Pain-1* alleles in tubers of the tetraploid cultivar ‘Satina’

The cultivar ‘Satina’ is tetraploid. Potentially four different *Pain-1* alleles can occur. Cloning and sequencing of ‘Satina’ cDNA resulted in two different alleles *Pain_SA* and *Pain_SN*.

In a pyrosequencing assay the alleles were analyzed on the basis of the A/C polymorphism of the associated SNP at cDNA position 1544 (Table 3.1.4). This SNP allows the comparison of the expression of *Pain_SA*, which exhibits the allele specific nucleotide A against the allelic fraction containing nucleotide C. The nucleotide C fraction can contain up to three different alleles, due to the ploidy state of the cultivated potatoes but is only represented by *Pain_SN*.

The pyrosequencing analysis was performed using cDNA from cold stored tubers and tuber genomic DNA to measure the dosages of the alleles (Figure 3.1.12).

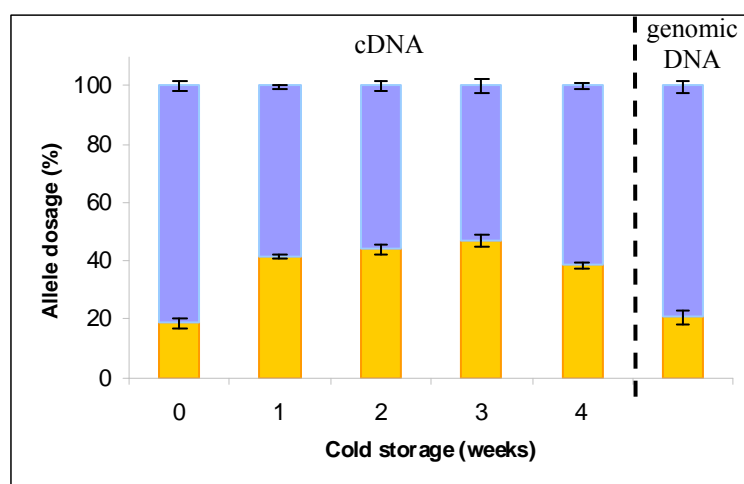


Figure 3.1.12: Pyrosequencing analysis of the alleles *Pain_SA* and *Pain_SN* of the cultivar ‘Satina’.

■ : *Pain_SA*; ■ : *Pain_SN*. For allele discrimination the primers Pyro_Pain_F/Pyro_Pain_RB (chapter 2, Table 2.1.4) were used. The primer Pyro_Pain_F was also used as sequencing primer. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of tubers. Percentages of the expression level of the cold stored samples are related to the genomic dosage of the alleles. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Standard deviations are derived from two biological replicates done in technical triplicates.

From ‘Satina’ two *Pain-1* alleles were identified. Pyrosequencing analysis of genomic tuber DNA revealed that the associated allele *Pain_SA* represents 25% of the potential four *Pain-1* alleles in the genotype ‘Satina’ (Figure 3.1.11). If the alleles were transcribed according to the genomic allele dosage, *Pain_SA* is expected to contribute 25% of the total transcripts and *Pain_SN* 75%. Comparison of allele dosage and allele transcription revealed that the alleles *Pain_SA* and *Pain_SN* differed in their abundance during tuber cold storage. The expression level of *Pain_SA* increased during tuber cold storage relative to *Pain_SN*. Maximal expression level was at $\approx 43\%$ in the 3rd week of cold storage.

Transferring the information about allelic distribution from pyrosequencing analysis to the total amount of invertase transcript levels from qRT-PCR resulted in detailed expression pattern during tuber cold storage (Figure 3.1.13).

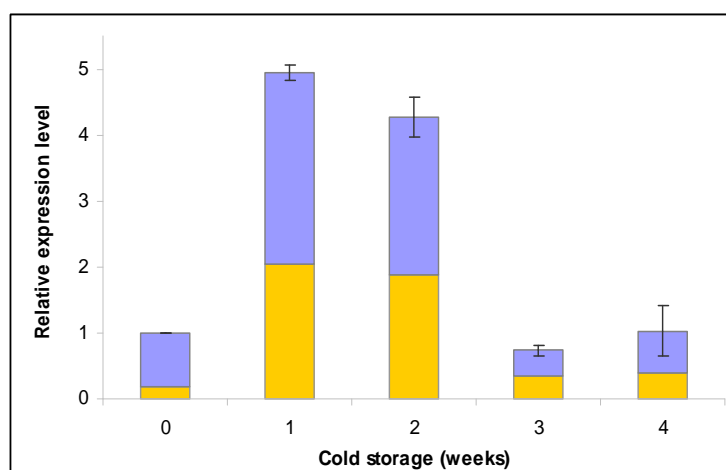


Figure 3.1.13: Overview of the expression pattern during tuber cold storage combining allelic distribution and total invertase transcript analysis of the cultivar ‘Satina’. ■: *Pain_SA*; ■: *Pain_SN*. The quantification of total invertase transcript was performed using the primers Pyro_Pain_F/Pyro_Pain_R (chapter 2, Table 2.1.9) in qRT-PCR analysis. Total transcript data were normalized with the reference gene *EF1a*. The relative expression level was calculated by defining transcript data of 0 weeks of cold storage as ‘1’. All other transcript data were related to this time point. Allelic distribution was calculated by relating the pyrosequencing data to the total transcript data. Standard deviations derived from two biological replicates done in technical duplicates. Standard deviations represent the variation within the replicates of the total transcripts, standard deviations for allele specific expression are shown in Figure 3.1.12.

qRT-PCR analysis showed an approximately four fold induction of total invertase transcripts after one week of tuber cold storage at 4°C (Figure 3.1.13). Transcript up-regulation declined in the 3rd and 4th week of cold storage to levels similar to samples not stored in the cold.

Expression analysis of the associated ‘Satina’ allele *Pain_SA*, which was found in one genomic dosage revealed an up-regulation up to 43% during tuber cold storage. Total invertase transcripts were induced up to four fold during the applied cold storage period. In

consequence tuber cold storage leads to a relative induction of the *Pain_SA* expression in contrast to the remaining allelic fraction in the tetraploid genotype ‘Satina’.

3.1.2.2.2 Expression pattern of *Pain-1* alleles in tubers of the tetraploid cultivar ‘Diana’

Referring to the cultivar tetraploidy, ‘Diana’ can consist of four possible *Pain-1* alleles. In this study three different alleles *Pain_DA*, *Pain_DN1*, and *Pain_DN2* were found. The expression of the three alleles was analyzed by tracing allele specific SNPs (Table 3.1.4; Table 3.1.13), which were used in pyrosequencing assays (Figure 3.1.14).

Table 3.1.13: Overview of allele specific SNPs analyzed by pyrosequencing.

Allele	SNP position	Allele specific SNP
<i>Pain_DA</i>	SNP 1544	<i>Pain_DA/Pain_DN1/Pain_DN2</i> A/C/C
<i>Pain_DN1</i>	SNP 1596	T/C/T
<i>Pain_DN2</i>	SNP 1574	A/A/T

SNP positions refer to cDNA sequence where ‘1’ represents the adenine of the start codon ATG. Primers used for pyrosequencing are listed in chapter 2, Table 2.1.4 (Pyro_Pain_F/Pyro_Pain_RB; sequencing primer Pyro_Pain_F). Allele specific SNPs are highlighted in bold capitals.

Pyrosequencing analysis was performed using cDNA from cold stored tubers and tuber genomic DNA to measure the dosages of the alleles (Figure 3.1.13).

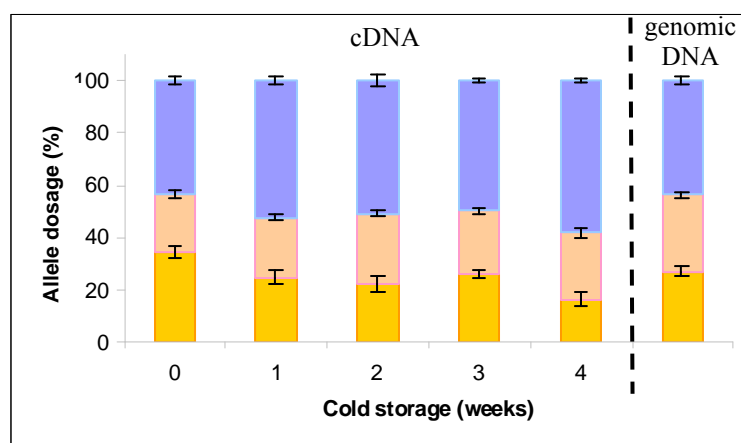


Figure 3.1.14: Pyrosequencing analysis of the alleles *Pain_DA*, *Pain_DN1*, and *Pain_DN2* of the cultivar ‘Diana’. ■: *Pain_DA*; ■: *Pain_DN1*; ■: *Pain_DN2*. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of tubers. Percentages of the expression level of the cold stored samples are related to the genomic dosage of the alleles. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Standard deviations are derived from two biological replicates done in technical triplicates.

Pyrosequencing analysis of genomic DNA showed that the ‘Diana’ alleles *Pain_DA* and *Pain_DN2* are present in simplex (25%), whilst *Pain_DN1* occurs in duplex (50%).

The allele *Pain_DN1* displays at SNP position 1596 the nucleotide C but it might be possible that this fraction, representing half of the allelic entity, contains more than one allele because

of tetraploidy. If the alleles were transcribed according to their allele dosage, *Pain_DA* and *Pain_DN2* are expected to contribute 25% of the total transcripts and *Pain_DN1* 50%.

The expression of the associated allele *Pain_DA* changed in tubers during cold treatment from 38% before storage to a level of 18% in tubers stored in the cold for four weeks. *Pain_DN1* was the most prevalent allele of the cultivar ‘Diana’ and was expressed from approximately 40% (0 weeks) up to 60% (4 weeks). The expression level of the third allele *Pain_DN2* did not show strong variation during tuber cold storage. The level of expression remained stable between 20 and 25%, and, therefore, similar to the genomic dosage.

Relating the pyrosequencing data to the total amounts of invertase transcripts obtained by qRT-PCR resulted in detailed information about invertase expression during tuber cold storage (Figure 3.1.15).

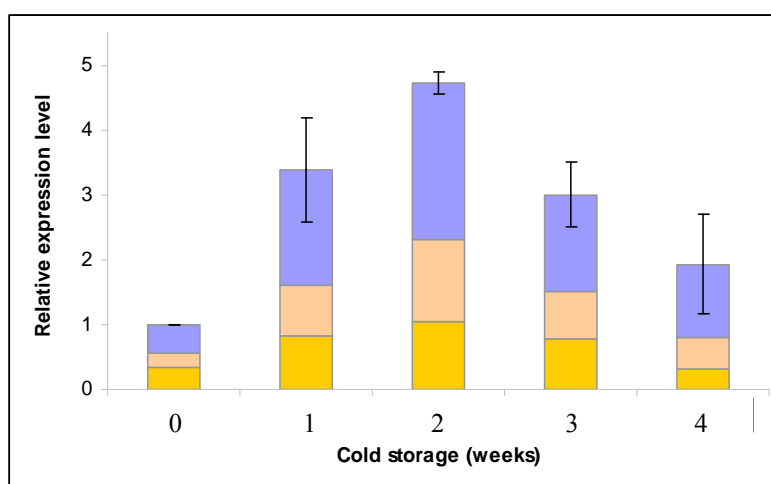


Figure 3.1.15: Overview of the expression pattern during tuber cold storage combining allelic distribution and total invertase transcript analysis of the cultivar ‘Diana’. ■: *Pain_DA*; ■: *Pain_DN1*; ■: *Pain_DN2*.

The quantification of total invertase transcript was performed using the primers Pyro_Pain_F/Pyro_Pain_R (chapter 2, Table 2.1.9) in qRT-PCR analysis. Total transcript data were normalized with the reference gene *EF1α* (section 2, Table 2.1.9). The relative expression level was calculated by defining transcript data of 0 weeks of cold storage as ‘1’. All other transcript data were related to this time point. Allelic distribution was calculated by relating the pyrosequencing data to the total transcript data. Standard deviations derived from two biological replicates done in technical duplicates. Standard deviations represent the variation within the replicates of the total transcripts, standard deviations for allele specific expression are shown in Figure 3.1.14.

qRT-PCR analysis showed an approximately 2.5 fold induction of total invertase transcript after one week of tuber cold storage (Figure 3.1.15). The transcript level increased in the 2nd week of cold storage up to about four fold compared to the control samples (no cold storage). In the 3rd and 4th week of storage the relative expression level declined but remained higher compared to levels in tubers not stored in the cold.

The expression of ‘Diana’ alleles during cold storage differed not strongly compared to their genomic dosages. Total invertase transcripts were induced up to three fold after the 1st and 2nd week of tuber cold storage.

3.1.2.2.3 Expression pattern of *Pain-1* alleles in tubers of the tetraploid cultivar ‘Theresa’

From the tetraploid cultivar ‘Theresa’ two different alleles *Pain_TN1* and *Pain_TN2* were obtained by cDNA cloning and sequencing. These two alleles were analyzed by means of the discriminative SNP at cDNA position 612 by pyrosequencing analysis (Figure 3.1.16). *Pain_TN1* has nucleotide A at SNP position 612, whilst *Pain_TN2* has nucleotide G (Table 3.1.4).

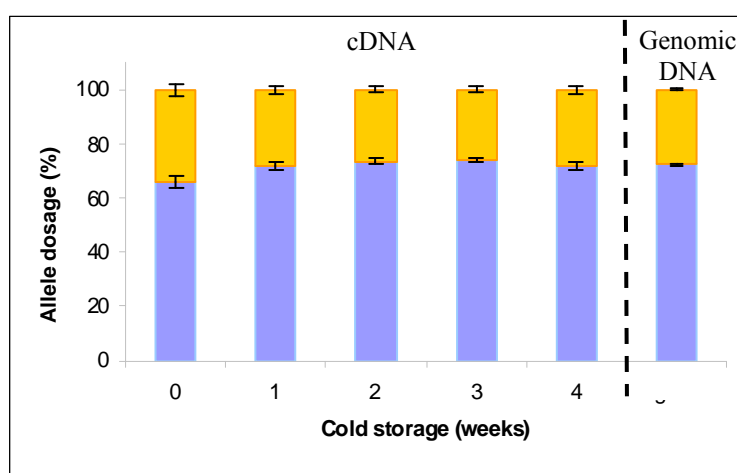


Figure 3.1.16: Pyrosequencing analysis of the alleles *Pain_TN1* and *Pain_TN2* of the cultivar ‘Theresa’. ■: *Pain_TN2*; ■: *Pain_TN1*. For allele discrimination the primers Pyro_PainTher_F/Pyro_PainTherRB (chapter 2, Table 2.1.4) were used. The primer Pyro_PainTheSeq was used as sequencing primer. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of tubers. Percentages of the expression level of the cold stored samples are related to the genomic dosage of the alleles. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Standard deviations are derived from two biological replicates done in technical triplicates.

Pyrosequencing analysis of genomic DNA showed that the allele *Pain_TN2* is present in simplex (25%) and the allele *Pain_TN1* in triplex (75%). If the alleles were transcribed according to the allele dosage, *Pain_TN1* is expected to contribute 75% of the total transcripts, and *Pain_TN2* 25%. Both alleles showed little changes in their presence in cDNA samples and their expression remained at similar levels during tuber cold treatment.

The expression level of *Pain_TN2* changed marginal during tuber cold storage compared with samples not stored at 4°C. *Pain_TN1* was the prevalent allele during tuber cold storage.

Referring to ‘Theresa’s’ tetraploidy, it is possible that the triplex fraction represented by *Pain_TN1* consists of more than one allele.

Pyrosequencing analysis gave information of allele expression during tuber cold storage. Transferring this allelic pattern to the total amount of invertase transcripts led to a detailed overview of *Pain-1* allele expression (Figure 3.1.17).

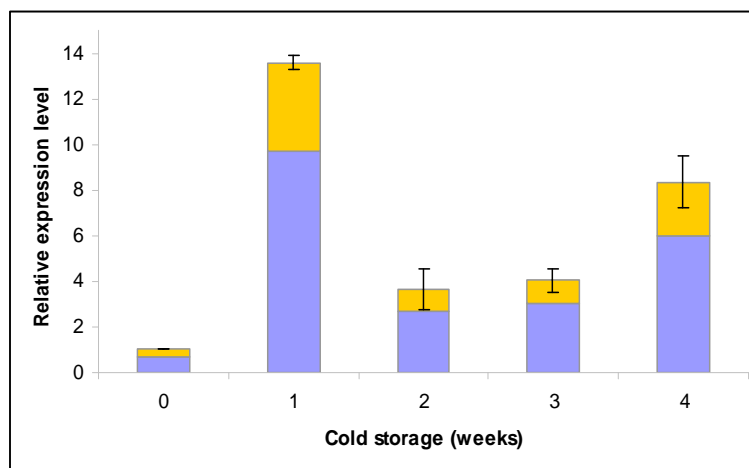


Figure 3.1.17: Overview of the expression pattern during tuber cold storage combining allelic distribution and total invertase transcript analysis of the cultivar ‘Theresa’. ■: *Pain-TN2*; ■: *Pain-TN1*. The quantification of total invertase transcript was performed using the primers Pyro_Pain_F/Pyro_Pain_R (chapter 2, Table 2.1.9) in qRT-PCR analysis. Total transcript data were normalized with the reference gene *EF1α* (chapter 2, Table 2.1.9). The relative expression level was calculated by defining transcript data of 0 weeks of cold storage as ‘1’. All other transcript data were related to this time point. Allelic distribution was calculated by relating the pyrosequencing data to the total transcript data. Standard deviations derived from three biological replicates done in technical duplicates. Standard deviations represent the variation within the replicates of the total transcripts, standard deviations for allele specific expression are shown in Figure 3.1.16.

qRT-PCR analysis showed an about 12 fold induction of total invertase transcript after the 1st week of tuber cold storage (Figure 3.1.17). Transcript up-regulation declined in the 2nd and 3rd week of cold storage to levels remaining higher than in control tubers (no cold storage). In the 4th week of cold treatment invertase transcripts increased again up to seven fold compared to control samples.

Expression analysis of both alleles of the cultivar ‘Theresa’ revealed no relevant changes of allele distribution in cold stored tubers. An intense up-regulation of total invertase transcripts was observed after the 1st week of tuber cold storage. The cultivar ‘Theresa’ is the only genotype analyzed, where total invertase transcripts increased again after four weeks at 4°C storage.

3.1.2.2.4 Expression pattern of *Pain-1* alleles in tubers of the diploid genotype P18

The differentiation between the two P18 alleles *Pain-P18N1* and *Pain-P18N2* relied on the C/A polymorphism of SNP at cDNA position 1544 using pyrosequencing analysis (Figure 3.1.18). The allele *Pain-P18N1* exhibits at SNP 1544 the nucleotide A, whilst allele *Pain-P18N2* consists of the nucleotide C (Table 3.1.5).

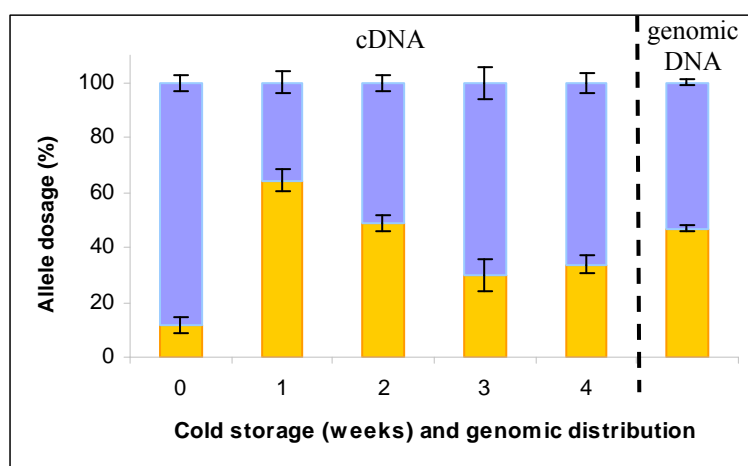


Figure 3.1.18: Pyrosequencing analysis of the alleles *Pain_P18N1* and *Pain_P18N2* of the genotype P18. ■: *Pain_P18N1*; ■: *Pain_P18N2*. For allele discrimination the primers Pyro Pyro_Pain_F/Pyro_Pain_RB (chapter 2, Table 2.1.4) were used. The primer Pyro_Pain_F was also used as sequencing primer. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of tubers. Percentages of the expression level of the cold stored samples are related to the genomic dosage of the alleles. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Standard deviations are derived from two biological replicates done in technical triplicates.

Pyrosequencing analysis of genomic DNA showed that both P18 alleles are present in duplex (50%) as expected for a heterozygous diploid genotype. Pyrosequencing analysis of cDNA samples showed a strong increase of allele *Pain_P18N1* after one week of tuber cold storage (Figure 3.1.18). Expression level was 50% higher than in tubers that were not stored in the cold. *Pain_P18N1* up-regulation declined in the 2nd, 3rd, and 4th week of cold storage to level ranging from about 30 to 50% of total expression.

Analyzing the relative expression level of P18 allele transcripts by transferring pyrosequencing data to qRT-PCR analysis showed that expression pattern changed during tuber cold storage (Figure 3.1.19).

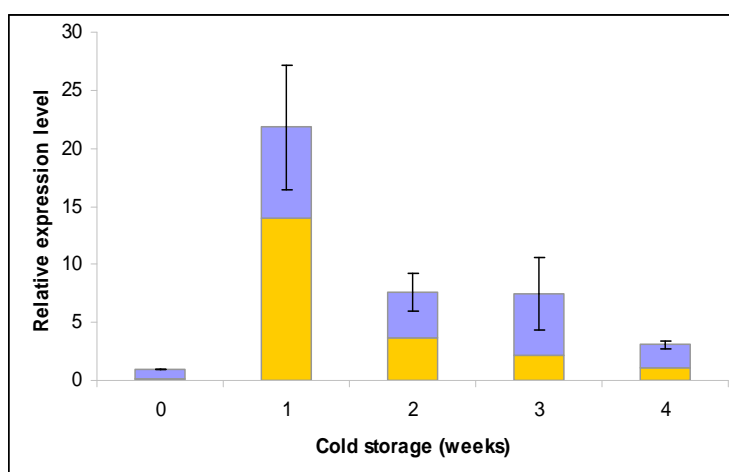


Figure 3.1.19: Overview of the expression pattern during tuber cold storage combining allelic distribution and total invertase transcript analysis of the potato genotype P18. ■: *Pain_P18N1*; ■: *Pain_P18N2*. The quantification of total invertase transcript was performed using the primers Pyro_Pain_F/Pyro_Pain_R (chapter 2, Table 2.1.9) in qRT-PCR analysis. Total transcript data were normalized with the reference gene *EF1α* (chapter 2, Table 2.1.9). The relative expression level was calculated by defining transcript data of 0 weeks of cold storage as '1'. All other transcript data were related to this time point. Allelic distribution was calculated by relating the pyrosequencing data to the total transcript data. Standard deviations derived from two biological replicates done in technical duplicates. Standard deviations represent the variation within the replicates of the total transcripts, standard deviations for allele specific expression are shown in Figure 3.1.18.

qRT-PCR analysis demonstrated that total invertase transcripts were strongly induced after one week of tuber cold storage at 4°C (Figure 3.1.19). Transcript up-regulation occurred up to 20 fold compared to control tubers (0 weeks). The relative expression level showed an about eight fold induction in the 2nd and 3rd week and decreased toward the end of cold storage.

Expression analysis of P18 invertase transcripts revealed an up-regulation of allele *Pain_P18N1* up to about 50% as well as an intense induction of total transcripts after one week of tuber cold storage. The diploid genotype P18 showed the highest accumulation of invertase transcripts (up to ≈20 fold) due to cold storage compared to control samples of all tested potato genotypes. Comparing the expression levels of the alleles *Pain_P18N1* and *Pain_DA*, which are identical at amino acid level, showed that expression of *Pain_DA* in contrast to *Pain_P18N1* was not effected by tuber cold storage (Figure 3.1.14). *Pain_DA* was found to be associated with better potato chips quality.

3.1.2.2.5 Expression pattern of *Pain-1* alleles in tubers of the diploid genotype P40

Pyrosequencing analysis of the P40 allele specific SNP at cDNA position 1267 were used to separate the alleles *Pain_P40N1* and *Pain_P40N2* (Figure 3.1.20). *Pain_P40N1* is characterized by adenine at SNP position 1267, whilst *Pain_P40N2* shows guanine (Table 3.1.5).

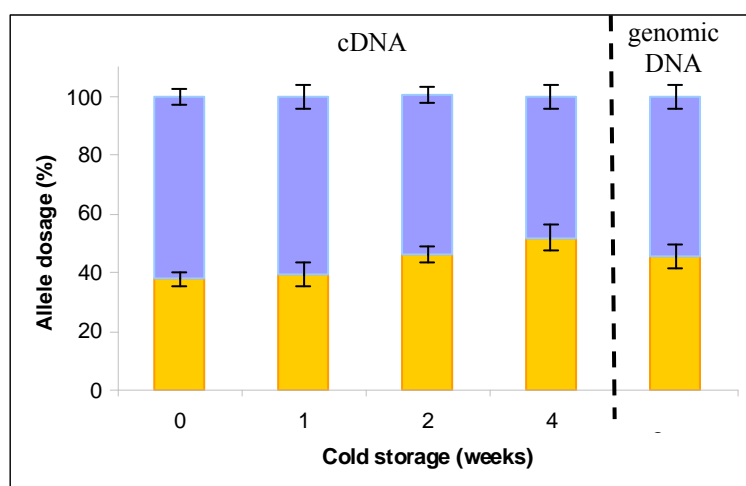


Figure 3.1.20: Pyrosequencing analysis of the alleles *Pain_P40N1* and *Pain_P40N2* of the genotype P40. ■ : *Pain_P40N1*; ■ : *Pain_P40N2*. For allele discrimination the primers Pyro_PainP40_F/Pyro_PainP40_RB were used. The primer Pyro_PainP40_Seq was used as sequencing primer (chapter 2, Table 2.1.4). In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of tubers. Percentages of the expression level of the cold stored samples are related to the genomic dosage of the alleles. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Standard deviations are derived from two biological replicates done in technical triplicates.

Pyrosequencing analysis of genomic DNA showed that both P 40 alleles are present in duplex (50%) as expected for a heterozygous diploid genotype. Pyrosequencing analysis of cDNA samples revealed a slight but continuous increase of allele *Pain_P40N1* during tuber cold storage (Figure 3.1.20). Expression levels ranged from about 38% in control samples up to 50% in tubers stored in the cold for four weeks. Tuber tissues stored for three weeks were not analyzed in pyrosequencing assay due to sample limitation.

Relating the pyrosequencing data to the total amounts of invertase transcripts obtained by qRT-PCR resulted in detailed information about invertase expression during tuber cold storage (Figure 3.1.21).

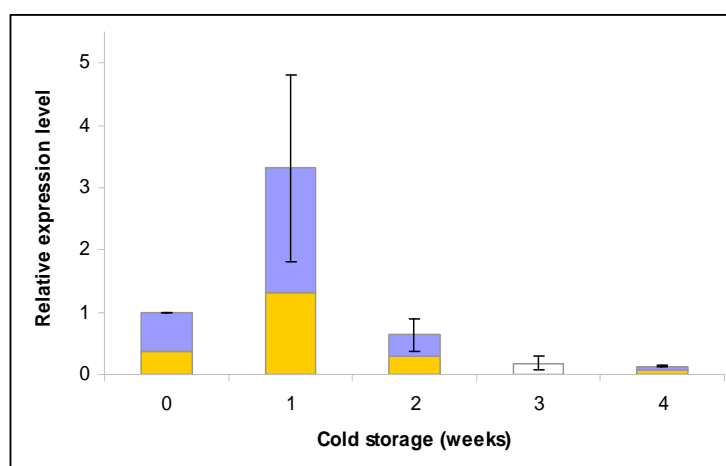


Figure 3.1.21: Overview of the expression pattern during tuber cold storage combining allelic distribution and total invertase transcript analysis of the potato genotype P40. ■ : *Pain_P40N1*; ■ : *Pain_P40N2*. The quantification of total invertase transcript was performed using the primers Pyro_Pain_F/Pyro_Pain_R (chapter 2, Table 2.1.9) in qRT-PCR analysis. Total transcript data were normalized with the reference gene *EF1α* (chapter 2, Table 2.1.9). The relative expression level was calculated by defining transcript data of 0 weeks of cold storage as '1'. All other transcript data were related to this time point. Allelic distribution was calculated by relating the pyrosequencing data to the total transcript data. The 3rd week of tuber cold storage was not analyzed by pyrosequencing due to samples limitations. The bar of total invertase transcripts, therefore, is left empty. Standard deviations derived from two biological replicates done in technical duplicates. Standard deviations represent the variation within the replicates of the total transcripts, standard deviations for allele specific expression are shown in Figure 3.1.20.

Monitoring the relative expression level of total invertase transcripts showed an about 2.5 fold induction after one week of tuber cold storage at 4°C (Figure 3.1.21). Transcript up-regulation declined in the 2nd, 3rd, and 4th week of cold storage to levels lower than in control tubers that were not stored in the cold.

Expression analysis of both P40 alleles revealed slight changes of allele distribution in cold stored tubers. Allele *Pain_P40N1* expression increased during cold storage from about 38% to 50%. Up-regulation of total invertase transcripts was observed after one week of cold treatment up to 2.5 fold. The relative expression level of total transcripts declined dramatically from the 2nd to the 4th week of tuber cold storage.

3.1.2.2 Functional complementation of the yeast invertase mutant *SUC2*

Plant invertases are functional in heterologous systems (FRIDMAN ET AL., 2004). Therefore, the model organism yeast (*Saccharomyces cerevisiae*) was used to analyze potato invertase alleles. The yeast invertase mutant *SUC2* lacks invertase activity and uses glucose as carbohydrate source. Transforming *SUC2* with *Pain-I* cDNA alleles resulted in yeast transformants that were able to grow on sucrose as sole carbohydrate source, indicating functional complementation of the *SUC2* mutation.

The *Pain-I* cDNA alleles listed below were used for complementation of *SUC2* (Figure 3.1.22) and subsequent analysis of invertase activity (section 3.1.2.3).


Genotype	Allele name	<i>SUC2</i> complementation
‘Satina’	<i>Pain_SA</i>	 <p>Representative yeast transformants complemented with invertase alleles of the cultivars ‘Satina’, ‘Diana’, and ‘Theresa’ were spotted on solid yeast minimal broth with 2% sucrose as carbon source. The wild type <i>FY 1479</i> was plated as positive control, whilst the invertase mutant <i>SUC2</i> was the negative control. Growth of all complemented yeast transformants was quantified (Figure 3.1.23).</p>
	<i>Pain_SN</i>	
‘Diana’	<i>Pain_DA</i>	
	<i>Pain_DN1</i>	
	<i>Pain_DN2</i>	
‘Theresa’	<i>Pain_TN1</i>	
	<i>Pain_TN2</i>	
P18	<i>Pain_P18N1</i>	
	<i>Pain_P18N2</i>	
P40	<i>Pain_P40N1</i>	
	<i>Pain_P40N2</i>	
P54	<i>Pain_P54N</i>	

Figure 3.1.22: cDNA alleles used for *SUC2* complementation and *SUC2* transformants on solid yeast minimal media.

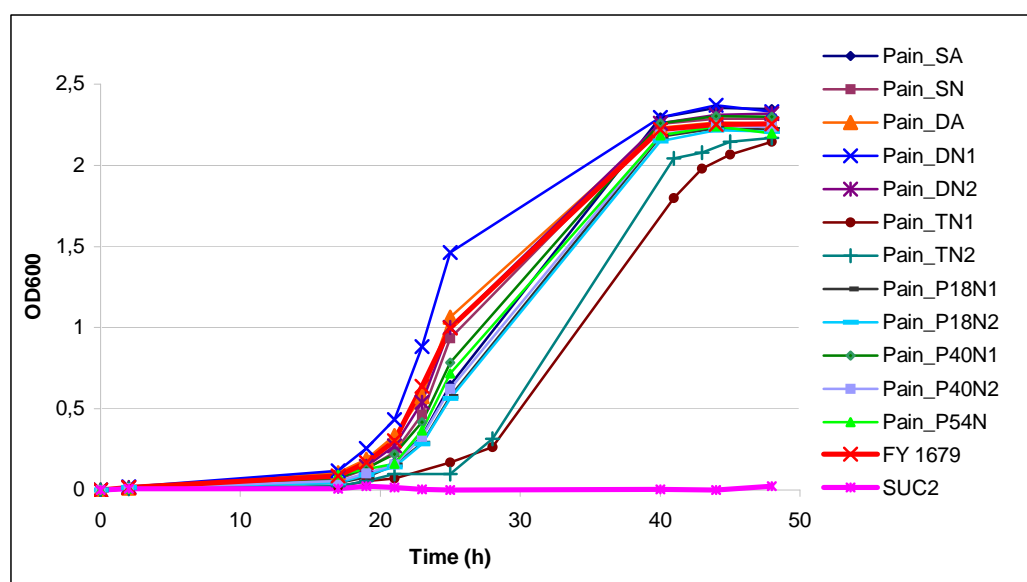


Figure 3.1.23: Growth of complemented *SUC2* transformants. Yeast reference strain *FY 1679* (red), yeast invertase mutant strain *SUC2* (pink), and *Pain-I* transformants were grown in yeast minimal media with 2% sucrose as the sole carbon source. OD600 is plotted against growth time in hours. OD600 values represent the means of two replicates. Standard deviations were less than 20% of mean.

All *SUC2* invertase transformants exhibited substantial growth on sucrose. ‘Satina’ and ‘Diana’ alleles and alleles of the diploid genotypes P18, P40, and P54 complemented the invertase deficiency of the yeast strain better than the alleles of ‘Theresa’. However, it was not possible to correlate the altered complementation efficiency to quantitative invertase protein differences using immunoblot studies (section 3.1.2.4; Figure 3.1.25).

3.1.2.3 Biochemical characterization of *Pain-1* alleles

Putative 3D-models of allelic *Pain-1* molecules (3.1.1.3) indicated structural and electrostatic differences between the alleles, which could cause functional differences. To test, whether structural characteristics might influence enzymatic activity of *Pain-1* invertase alleles, biochemical characterization was performed.

From yeast *SUC2* transformants complemented with *Pain-1* cDNA alleles total protein was extracted. To test soluble acid invertase activity, a modified protocol based on ZRENNER ET AL. (1995) was used. Invertase assays were carried out at 30°C. Additionally, *Pain-1* alleles were assayed at 4°C. The analysis at 4°C was performed to study possible differences of enzyme kinetics due to allelic amino acid composition in response to cold storage conditions. It was previously reported that cold storage influences transcriptional changes regarding vacuolar invertases (ZRENNER ET AL., 1996; ZHOU ET AL., 2004; BAGNARESI ET AL., 2008), but no study investigated the effect of low temperatures on the enzyme activity itself. The *Pain-1* alleles displayed in Table 3.1.14 were used for biochemical characterization at 30°C and 4°C.

Table 3.1.14: *Pain-1* alleles used for biochemical characterization.

Tetraploid genotypes	Allele	Diploid genotypes	Allele
‘Satina’	<i>Pain_SA</i>	P18	<i>Pain_P18N1</i>
	<i>Pain_SN</i>		<i>Pain_P18N2</i>
‘Diana’	<i>Pain_DA</i>	P40	<i>Pain_P40N1</i>
	<i>Pain_DN1</i>		<i>Pain_P40N2</i>
	<i>Pain_DN2</i>	P54	<i>Pain_P54N</i>
‘Theresa’	<i>Pain_TN1</i>		
	<i>Pain_TN2</i>		

The biochemical parameters Michaelis constant (K_m) and the maximal velocity (v_{max}) of invertase reaction were determined.

Results of the biochemical analysis at 30°C and 4°C are shown in Table 3.1.15.

Table 3.1.15: K_m (mM) and v_{max} (mmol/h*mg protein⁻¹) of *Pain-I* invertase alleles at 30°C and 4°C.

Genotype	Allele	K_m (30°C)	K_m (4°C)	v_{max} (30°C)	v_{max} (4°C)
‘Satina’	<i>Pain_SA</i>	22.1±1.4	2.3±0.3	4.7±1.0	1.7±0.1
	<i>Pain_SN</i>	23.0±1.4	3.5±0.3	4.1±1.0	1.5±0.1
‘Diana’	<i>Pain_DA</i>	19.9±1.4	4.0±0.3	12.4±0.8	2.8±0.1
	<i>Pain_DN1</i>	19.6±1.4	2.7±0.3	11.2±0.8	2.7±0.1
	<i>Pain_DN2</i>	15.6±1.4	3.7±0.3	6.2±1.0	2.2±0.1
‘Theresa’	<i>Pain_TN1</i>	19.8±1.4	3.0±0.3	3.4±1.0	1.5±0.1
	<i>Pain_TN2</i>	21.6±1.4	na	2.0±1.4	na
P18	<i>Pain_P18N1</i>	20.4±1.4	2.7±0.3	5.5±1.0	2.1±0.1
	<i>Pain_P18N2</i>	16.9±1.4	4.7±0.4	2.7±1.0	1.9±0.1
P40	<i>Pain_P40N1</i>	17.2±1.4	3.4±0.3	5.8±1.0	2.1±0.1
	<i>Pain_P40N2</i>	14.9±1.4	4.4±0.3	6.8±1.0	2.6±0.1
P54	<i>Pain_P54N</i>	19.1±1.4	2.7±0.4	3.7±1.0	1.6±0.1
<i>FY 1679</i>		21.0±1.6	21.1±2.5	23.9±4.5	7.7±0.9

Standard deviations are derived from three biological replicates for the associated alleles *Pain_SA* and *Pain_DA*, and the wild type reference strain *FY 1679*, and from two biological replicates for the other alleles done in technical replicates to obtain six measurements. To make assays of different invertase isoforms comparable, the yeast reference strain *FY 1679* was used as positive control.

Tables 3.1.16, 3.1.17, 3.1.18 and 3.1.19 summarize the significance values for differences between the K_m and v_{max} values measured for the *Pain-I* alleles at 30°C and 4°C. The statistical calculation was performed by Benjamin Stich, MPIZ/Köln.

Table 3.1.16: Overview of statistical significance levels of K_m values from the *Pain-I* invertase alleles at 30°C.

Allele	<i>Pain_SA</i>	<i>Pain_SN</i>	<i>Pain_DA</i>	<i>Pain_DN1</i>	<i>Pain_DN2</i>	<i>Pain_TN1</i>	<i>Pain_TN2</i>	<i>Pain_P18N1</i>	<i>Pain_P18N2</i>	<i>Pain_P40N1</i>	<i>Pain_P40N2</i>	<i>Pain_P54N</i>
<i>Pain_SA</i>	---	0.63	0.28	0.22	0.0058	0.26	0.79	0.39	0.019	0.028	0.0029	0.15
<i>Pain_SN</i>	---	---	0.13	0.10	0.0024	0.12	0.47	0.20	0.008	0.011	0.0012	0.067
<i>Pain_DA</i>	---	---	---	0.89	0.048	0.98	0.40	0.80	0.15	0.20	0.024	0.71
<i>Pain_DN1</i>	---	---	---	---	0.06	0.91	0.33	0.70	0.19	0.25	0.03	0.81
<i>Pain_DN2</i>	---	---	---	---	---	0.051	0.009	0.03	0.53	0.42	0.71	0.095
<i>Pain_TN1</i>	---	---	---	---	---	---	0.38	0.78	0.16	0.21	0.025	0.73
<i>Pain_TN2</i>	---	---	---	---	---	---	---	0.55	0.032	0.045	0.0047	0.23
<i>Pain_P18N1</i>	---	---	---	---	---	---	---	---	0.09	0.13	0.015	0.53
<i>Pain_P18N2</i>	---	---	---	---	---	---	---	---	---	0.86	0.32	0.27
<i>Pain_P40N1</i>	---	---	---	---	---	---	---	---	---	---	0.25	0.35
<i>Pain_P40N2</i>	---	---	---	---	---	---	---	---	---	---	---	0.048
<i>Pain_P54N</i>	---	---	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

Table 3.1.17: Overview of statistical significance levels of K_m values from the *Pain-1* invertase alleles at 4°C.

Allele	<i>Pain SA</i>	<i>Pain SN</i>	<i>Pain DA</i>	<i>Pain DN1</i>	<i>Pain DN2</i>	<i>Pain TN1</i>	<i>Pain TN2</i>	<i>Pain P18N1</i>	<i>Pain P18N2</i>	<i>Pain P40N1</i>	<i>Pain P40N2</i>	<i>Pain P54N</i>
<i>Pain SA</i>	---	0.014	0.0019	0.36	0.0078	0.11	na	0.298	0.0003	0.023	0.0004	0.34
<i>Pain SN</i>	---	---	0.24	0.080	0.65	0.33	na	0.1	0.026	0.798	0.06	0.14
<i>Pain DA</i>	---	---	---	0.0099	0.47	0.053	na	0.013	0.21	0.16	0.44	0.022
<i>Pain DN1</i>	---	---	---	---	0.042	0.43	na	0.89	0.0012	0.13	0.0021	0.89
<i>Pain DN2</i>	---	---	---	---	---	0.18	na	0.052	0.066	0.49	0.15	0.079
<i>Pain TN1</i>	---	---	---	---	---	---	na	0.51	0.0058	0.46	0.012	0.57
<i>Pain TN2</i>	---	---	---	---	---	---	---	na	na	na	na	na
<i>Pain P18N1</i>	---	---	---	---	---	---	---	---	0.0014	0.16	0.0027	0.98
<i>Pain P18N2</i>	---	---	---	---	---	---	---	---	---	0.017	0.59	0.0028
<i>Pain P40N1</i>	---	---	---	---	---	---	---	---	---	---	0.038	0.2085
<i>Pain P40N2</i>	---	---	---	---	---	---	---	---	---	---	---	0.0055
<i>Pain P54N</i>	---	---	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

Table 3.1.18: Overview of statistical significance levels of v_{max} values from the *Pain-1* invertase alleles at 30°C.

Allele	<i>Pain SA</i>	<i>Pain SN</i>	<i>Pain DA</i>	<i>Pain DN1</i>	<i>Pain DN2</i>	<i>Pain TN1</i>	<i>Pain TN2</i>	<i>Pain P18N1</i>	<i>Pain P18N2</i>	<i>Pain P40N1</i>	<i>Pain P40N2</i>	<i>Pain P54N</i>
<i>Pain SA</i>	---	0.696	5.80E-05	0.0003	0.29	0.38	0.15	0.58	0.177	0.43	0.15	0.52
<i>Pain SN</i>	---	---	3.01E-05	0.0001	0.16	0.62	0.25	0.35	0.32	0.25	0.077	0.79
<i>Pain DA</i>	---	---	---	0.32	0.0004	1.35E-05	3.11E-05	0.0002	6.2E-06	0.0002	0.0009	1.98E-05
<i>Pain DN1</i>	---	---	---	---	0.0021	5.26E-05	9.68E-05	0.0007	2.28E-05	0.0012	0.0049	7.98E-05
<i>Pain DN2</i>	---	---	---	---	---	0.0653	0.03	0.599	0.026	0.77	0.68	0.1
<i>Pain TN1</i>	---	---	---	---	---	---	0.44	0.16	0.61	0.11	0.03	0.81
<i>Pain TN2</i>	---	---	---	---	---	---	---	0.067	0.72	0.047	0.016	0.34
<i>Pain P18N1</i>	---	---	---	---	---	---	---	---	0.068	0.813	0.356	0.249
<i>Pain P18N2</i>	---	---	---	---	---	---	---	---	---	0.044	0.012	0.46
<i>Pain P40N1</i>	---	---	---	---	---	---	---	---	---	---	0.49	0.16
<i>Pain P40N2</i>	---	---	---	---	---	---	---	---	---	---	---	0.048
<i>Pain P54N</i>	---	---	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

Table 3.1.19: Overview of statistical significance levels of v_{\max} values from the *Pain-1* invertase alleles at 4°C.

Allele	<i>Pain</i> <i>SA</i>	<i>Pain</i> <i>SN</i>	<i>Pain</i> <i>DA</i>	<i>Pain</i> <i>DN1</i>	<i>Pain</i> <i>DN2</i>	<i>Pain</i> <i>TN1</i>	<i>Pain</i> <i>TN2</i>	<i>Pain</i> <i>P18N1</i>	<i>Pain</i> <i>P18N2</i>	<i>Pain</i> <i>P40N1</i>	<i>Pain</i> <i>P40N2</i>	<i>Pain</i> <i>P54N</i>
<i>Pain</i> <i>SA</i>	---	0.06	4.05 E-06	8.19 E-06	0.0093	0.22	na	0.013	0.25	0.013	3.20 E-05	0.55
<i>Pain</i> <i>SN</i>	---	---	3.74 E-07	6.11 E-07	0.0003	0.5	na	0.0003	0.0097	0.0003	2.18 E-06	0.23
<i>Pain</i> <i>DA</i>	---	---	---	0.41	0.0006	1.22 E-06	na	0.0003	5.86 E-05	0.0003	0.2	4.80 E-06
<i>Pain</i> <i>DN1</i>	---	---	---	---	0.0019	2.20E- 06	na	0.0008	0.0002	0.0008	0.58	9.54 E-06
<i>Pain</i> <i>DN2</i>	---	---	---	---	---	0.0012	na	0.78	0.12	0.77	0.0073	0.0052
<i>Pain</i> <i>TN1</i>	---	---	---	---	---	---	na	0.0015	0.037	0.0015	7.64 E-06	0.56
<i>Pain</i> <i>TN2</i>	---	---	---	---	---	---	---	na	na	na	na	na
<i>Pain</i> <i>P18N1</i>	---	---	---	---	---	---	---	---	0.17	0.99	0.0034	0.0069
<i>Pain</i> <i>P18N2</i>	---	---	---	---	---	---	---	---	---	0.18	0.0005	0.12
<i>Pain</i> <i>P40N1</i>	---	---	---	---	---	---	---	---	---	---	0.0034	0.0067
<i>Pain</i> <i>P40N2</i>	---	---	---	---	---	---	---	---	---	---	---	3.14 E-05
<i>Pain</i> <i>P54N</i>	---	---	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

The 30°C assay of *Pain-1* alleles showed that the substrate affinity ranged between 15mM of the allele *Pain_P40N2* and 23mM of the allele *Pain_SN*. Maximal velocities of the alleles varied between 2mmol*h⁻¹*mg protein⁻¹ in the case of the allele *Pain_TN2* and 12mmol*h⁻¹*mg protein⁻¹ for the allele *Pain_DA*. The enzymatic characteristics of the analyzed alleles from the cultivar ‘Satina’ *Pain_SA* and *Pain_SN* showed no significant differences. In the K_m and v_{\max} values of the cultivar ‘Diana’ differences were displayed in respect to the allele *Pain_DN2*. *Pain_DN2* showed the highest substrate affinity with a K_m of approximately 16mM, and the slowest substrate conversion with a v_{\max} of around 6mmol*h⁻¹*mg protein⁻¹. The other two analyzed alleles *Pain_DA* and *Pain_DN1* displayed substrate affinities of approximately 20mM and maximal velocities of around 12mmol*h⁻¹*mg protein⁻¹. Comparison of the associated *Pain-1* alleles *Pain_SA* and *Pain_DA*, which are not identical at amino acid level, revealed no significant differences regarding their affinity to sucrose, but showed strong differences in the rate of sucrose conversion. The allele *Pain_DA* converts sucrose approximately 2.5 times faster than the allele *Pain_SA*.

Looking at the other analyzed *Pain-1* alleles, the two ‘Theresa’ alleles showed similar enzymatic characteristics and did not differ significantly. K_m values varied from 20 to 21mM, and v_{\max} values ranged from 2 to 3mmol*h⁻¹*mg protein⁻¹. Biochemical analysis of the alleles

from the diploid potato genotypes P18 and P40 did not display any significant K_m and v_{max} differences. K_m and v_{max} values of *Pain_P18N1* and *Pain_P18N2* ranged from 17 to 20mM, and 3 to 5.5mmol*h⁻¹*mg protein⁻¹, respectively. The alleles *Pain_P40N1* and *Pain_P40N2* displayed substrate affinities between 15 and 17mM, and v_{max} values between 6 and 7mmol*h⁻¹*mg protein⁻¹.

Biochemical analysis of the enzymatic characteristics of the *Pain-I* alleles at 4°C showed a dramatic increase in the enzyme's affinity to sucrose. The K_m values decreased approximately 5.5 times compared to K_m values measured at 30°C. Also the maximal velocities of the alleles were affected at 4°C. By trend v_{max} values were around 2mmol*h⁻¹*mg protein⁻¹.

Additionally, biochemical characteristics were determined for two nucleotide variants of the 'Satina' allele *Pain_SN* to measure codon influences in the heterologous system yeast. The variants differed in one to three nucleotides to *Pain_SN* (Table 3.1.20), (Appendix A 3.1.25).

Table 3.1.20: SNPs of the nucleotide variants of the allele *Pain_SN*.

SNP position	<i>Pain_SN</i>	<i>Pain_SN</i> *	<i>Pain_SN</i> **
69	T	T	C
75	T	C	T
1050	C	T	C
1350	A	G	A

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. The nucleotide exchanges are synonymous, not causing an amino acid exchange.

It was shown that all nucleotide variants of *Pain_SN* displayed similar K_m and v_{max} values (Table 3.1.21), showing that a codon usage independent comparison of different alleles was possible.

Table 3.1.21: K_m (mM) and v_{max} (mmol/h*mg protein⁻¹) of *Pain-SN1* nucleotide variants at 30°C and 4°C.

Allele	K_m (30°C)	K_m (4°C)	v_{max} (30°C)	v_{max} (4°C)
<i>Pain SN</i>	23.0±1.4	3.5±0.3	4.1±1.0	1.5±0.1
<i>Pain SN</i> *	23.0±1.7	3.0±0.3	5.7±1.0	1.8±0.1
<i>Pain SN</i> **	24.5±1.5	2.5±0.4	7.8±1.4	1.7±0.1

Standard deviations are derived from two biological replicates done in technical triplicates.

The Tables 3.1.22 and 23 summarize the significance values for differences between the K_m and v_{max} values measured for the nucleotide variants of the allele *Pain_SN* at 30°C and 4°C. The statistical calculation was performed by Benjamin Stich, MPIZ/Köln.

Table 3.1.22: Overview of statistical significance levels of K_m values from the *Pain-SN1* nucleotide variants at 30°C and 4°C.

Allele	30°C			4°C		
	<i>Pain</i> <i>SN</i>	<i>Pain</i> <i>SN*</i>	<i>Pain</i> <i>SN**</i>	<i>Pain</i> <i>SN</i>	<i>Pain</i> <i>SN*</i>	<i>Pain</i> <i>SN**</i>
<i>Pain</i> <i>SN</i>	---	0.1	0.48	---	0.28	0.06
<i>Pain</i> <i>SN*</i>	---	---	0.53	---	---	0.35
<i>Pain</i> <i>SN**</i>	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

Table 3.1.23: Overview of statistical significance levels of v_{max} values from the *Pain-SN1* nucleotide variants at 30°C and 4°C.

Allele	30°C			4°C		
	<i>Pain</i> <i>SN</i>	<i>Pain</i> <i>SN*</i>	<i>Pain</i> <i>SN**</i>	<i>Pain</i> <i>SN</i>	<i>Pain</i> <i>SN*</i>	<i>Pain</i> <i>SN**</i>
<i>Pain</i> <i>SN</i>	---	0.27	0.048	---	0.04	0.084
<i>Pain</i> <i>SN*</i>	---	---	0.24	---	---	0.83
<i>Pain</i> <i>SN**</i>	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are orange coloured, no significant differences are described by P-values >0.05.

3.1.2.4 Western blot analysis

To analyze, whether the observed differences of invertase enzyme activity and yeast growth behaviour are not due to changes in protein quantity, immunoblot quantification was performed.

As a loading control the blotted membrane was stained with Ponceau S (Figure 3.1.24). The concentration of total yeast protein extract was the same within the loaded samples.

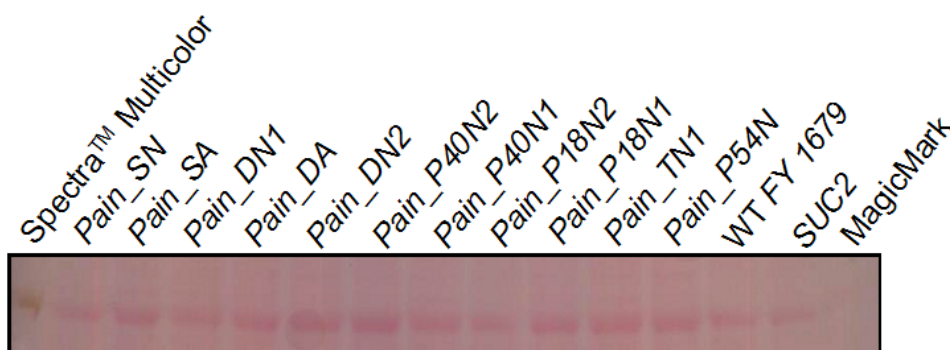


Figure 3.1.24: Ponceau S stained blot membrane. The corresponding alleles are given by their name above every lane. 15µg of total protein were loaded. A representative protein band out of the blotted membrane is shown as loading control.

From all tested invertase antibodies (chapter 2, Table 2.2.15), the antibody against a 58kDa vacuolar invertase of potato (BURCH ET AL., 1992) detected *Pain-1* protein most reliable and, therefore, was used in subsequent Western blot analysis. The invertase protein content of a set of allelic *Pain-1* yeast *SUC2* transformants was analyzed representatively (Figure 3.1.25).

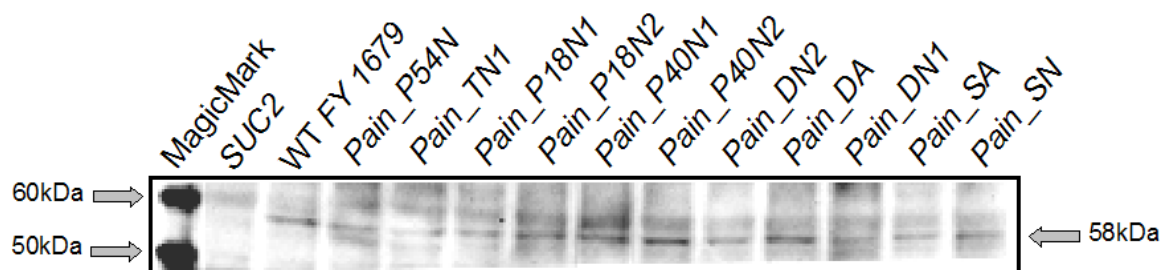


Figure 3.1.25: Western blot analysis of *Pain-1* alleles using an antibody against a 58kDa vacuolar invertase of potato. The corresponding alleles are given by their name above every lane. 15µg of total protein were loaded. Additionally, yeast invertase mutant *SUC2* and yeast reference strain WT *FY 1679* were blotted. Left arrows stand for size bands of the MagicMark Marker, the right arrow refers to the 58kDa band where the invertase protein was detected.

Western blot analysis showed that invertase protein content of *SUC2* transformants containing different *Pain-1* alleles did not differ significantly. Additionally, the invertase of the yeast reference strain WT *FY 1679* was detected by the potato antibody compared to the yeast invertase mutant *SUC2* where no protein was detectable.

3.2 The *Invap-b* locus on chromosome IX

The *Invap-b* locus was intensively studied and described by MADDISON ET AL., (1999). The locus consists of two invertase genes *invGE* and *invGF* linked in direct tandem repeat and has a size of approximately 8.6kb. Both genes exhibit a similar exon/intron structure composed of six exons and five introns. This exon/intron structure is identical to the two potato cell wall-bound invertases *pCD111* and *pCD141* on chromosome X (HEDLEY ET AL., 1993, 1994), and slightly different from the *Pain-1* invertase gene on chromosome III (ZHOU ET AL., 1994). Expression analysis showed *invGE* transcription in the leaf, stem, root, tuber, and floral tissue, whilst *invGF* expression is restricted to floral tissues (MADDISON ET AL., 1999).

The genes *invGE* and *invGF* encode cell wall-bound invertase isoforms. These acidic insoluble invertases act in a pH optimum range of 4.5 to 5.0 and are functional in the apoplast (TYMOWSKA-LALANNE & KREIS, 1998).

3.2.1 Structural characterization of the genes *invGE* and *invGF*

3.2.1.1 Identification of associated *invGE* and *invGF* alleles

The *Invap-b* locus maps to potato chromosome IX (CHEN ET AL., 2001) in a region associated with tuber quality trait,s where a QTL for potato tuber sugar content, *Sug9a*, was identified (MENÉNDEZ ET AL., 2002). In the latter study, the genes *invGE* and *invGF* were directly mapped in the mapping population used and showed linkage to the QTL *Sug9a*.

Single-strand conformation polymorphism (SSCP) analysis revealed an association of *invGE* and *invGF* alleles with starch and sugar content of potato tubers. SSCP fragments found to be associated with better potato chips quality were named *invGE-6f* and *invGF-4d*, respectively (LI ET AL., 2005). The occurrence of those fragments in the genotypes was highly correlated indicating that *invGE-6f* and *invGF-4d* are in LD. The analysis of protein sequences of allelic *invGE-6f* fragments revealed a unique histidine instead of proline at amino acid position 368 (LI ET AL., 2005). *invGE-6f* fragments containing histidine at position 368 were associated with better potato chips quality. Histidine 368 is encoded by the cDNA SNP at nucleotide position 1103* (section 3.2.1.2, Table 3.2.3) where cytosine changes to adenine. Therefore, cloned *invGE* cDNA alleles were named based on the presence (A) or absence (N) of SNP 1103*.

Assigning cloned *invGF* alleles to the SSCP fragment *invGF-4d* and detecting the underlying SNPs are subject of ongoing investigations in the research project.

3.2.1.2 Molecular cloning of *invGE* and *invGF* invertase cDNA alleles

Regarding *invGE* and *invGF* expression patterns (MADDISON ET AL., 1999), molecular cloning of *invGE* alleles was performed using leaf and floral tissue, whilst *invGF* allele cloning was initially carried out with floral tissue. During this study *invGF* alleles were obtained also from leaf material demonstrating that the restriction of floral *invGF* expression is genotype dependent.

❖ Genotype dependent *invGF* expression in leaves

Following the approach to detect *invGF* transcripts as described by MADDISON ET AL. (1999), gene specific primers amplifying a small fragment of the gene from the tetraploid genotypes ‘Désirée’, ‘Saturna’, ‘Diana’, ‘Theresa’, and from the diploid genotypes P40 and P54 were designed. MADDISON ET AL. (1999) restricted the expression analysis consisting of a histochemical GUS assay of *invGF* expression in transgenic plants and RT-PCR from mature flowers, flower bud, source, and sink leaf, respectively to the genotypes ‘Désirée’ and ‘Saturna’. The *invGF* primers (chapter 2, Table 2.1.7) were selected spanning from exon I to exon III, which should generate a product of 402bp from cDNA. In contrast, any product generated from contaminating genomic DNA would include sequences from two intermediate introns and the mini-exon II, amounting to 704bp. RT-PCR with the latter primers using total RNA prepared from both mature flowers and leaves as template generated a product of the expected size (Figure 3.2.1) indicating expression of *invGF* in these organs.

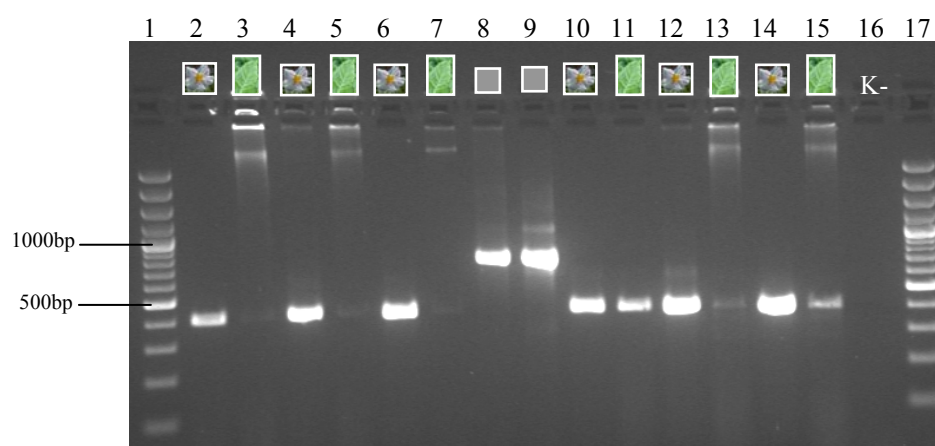


Figure 3.2.1: Amplification of a 402bp *invGF* fragment following the approach described by MADDISON ET AL. (1999). As template cDNA from leaves and flowers, and genomic DNA from leaves were used. cDNA template from flowers, cDNA template from leaves, genomic DNA from leaves. Lane 1: Fermentas ladder 100bp plus, lane 2: floral cDNA ‘Saturna’, lane 3: leaf cDNA ‘Saturna’, lane 4: floral cDNA ‘Désirée’, lane 5: leaf cDNA ‘Désirée’, lane 6: floral cDNA P40, lane 7: leaf cDNA P40, lane 8: genomic DNA P40, lane 9: genomic DNA ‘Diana’, lane 10: floral cDNA ‘Diana’, lane 11: leaf cDNA ‘Diana’, lane 12: floral cDNA ‘Theresa’, lane 13: leaf cDNA ‘Theresa’, lane 14: floral cDNA P54, lane 15: leaf cDNA P54, lane 16: negative control for leaf cDNA, lane 17: Fermentas ladder 100bp plus.

To obtain full-length *invGF* alleles, PCR amplification using full-length gene specific primers was accomplished. cDNA and genomic DNA of flowers and leaves from the genotypes ‘Satina’, ‘Diana’, ‘Theresa’, P18, P40, and P54 served as template. Additionally, full-length PCR was carried out for cDNA and genomic DNA from flowers and leaves of the cultivar ‘Saturna’ (Figure 3.2.2).

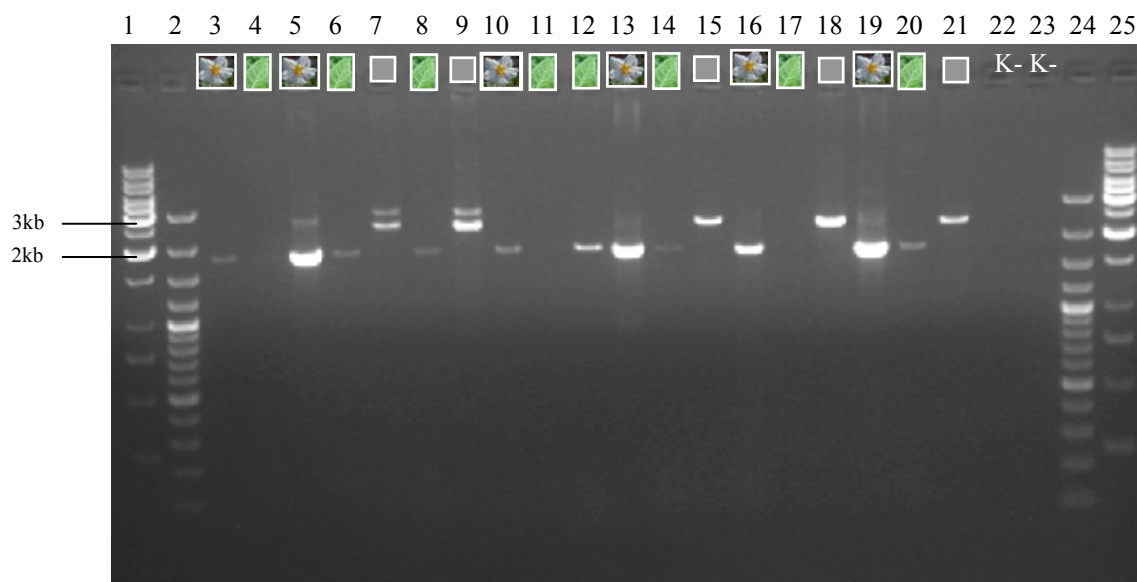


Figure 3.2.2: Full-length amplification of the gene *invGF* in different genotypes. As template cDNA from leaves and flowers, and genomic DNA from leaves were used. Full-length amplification was carried out using specific *invGF* primers (chapter 2, Table 2.1.2, *invGF*-F-fulgth/*invGF*-R-fulgth). A positive control was included using specific primers for the gene *pCD141* (chapter 2, Table 2.1.2, *CD141*f_l F/*CD141*f_l R) and cDNA from leaves to check cDNA integrity (lane 12). 🌸 cDNA template from flowers, 🌿 cDNA template from leaves, ■ genomic DNA from leaves. Lane1: 1kb Fermentas ladder, lane 2: Fermentas ladder 100bp plus, lane 3: floral cDNA ‘Satina’, lane 4: leaf cDNA ‘Satina’, lane 5: floral cDNA ‘Diana’, lane 6: leaf cDNA ‘Diana’, lane 7: genomic DNA ‘Diana’, lane 8: leaf cDNA ‘Theresa’, lane 9: genomic DNA ‘Theresa’, lane 10: floral cDNA ‘Saturna’, lane 11: leaf cDNA ‘Saturna’, lane 12: leaf cDNA ‘Saturna’ with *pCD141* specific primers as positive control, lane 13: floral cDNA P18, lane 14: leaf cDNA P18, lane 15: genomic DNA P18, lane 16: floral cDNA P40, lane 17: leaf cDNA P40, lane 18: genomic DNA P40, lane 19: floral cDNA P54, lane 20: leaf cDNA P54; lane 21: genomic DNA P54, lane 22: negative control for leaf cDNA, lane 23: negative control for genomic DNA, lane 24: Fermentas ladder 100bp plus, lane 25: 1kb Fermentas ladder.

Both PCRs with short and full-length primers revealed genotype depending *invGF* expression. *invGF* transcripts were detected in leaves of the genotypes ‘Diana’, ‘Theresa’, P18, and P54. The gene specific expression in the genotypes ‘Diana’ and P54 were stronger compared to ‘Theresa’ and P18. The genotype ‘Satina’ showed a very weak *invGF* expression (Figure 3.2.2 lane 4), No *invGF* transcripts were detectable in ‘Désirée’ (Figure 3.2.1 lane 5), ‘Saturna’ (Figure 3.2.1 lane 3, Figure 3.2.2 lane 11), and P40 (Figure 3.2.1 lane 7, Figure 3.2.2 lane 17).

Cloning and sequencing of the entire RT-PCR products confirmed their identity to and, therefore, its origin from *invGF*.

❖ *invGE* and *invGF* cDNA alleles

Using full-length gene specific primers, cDNA invertase alleles of *invGE* and *invGF* were cloned and sequenced from the three tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’ and from three diploid potato genotypes P18, P40, and P54.

All tetraploid genotypes harbour the associated SSCP fragments of the genes *invGE* and *invGF* (LI ET AL., 2005; Table 3.2.1) and, therefore, were chosen in this study. The diploid genotypes were included because they were parents of the QTL analysis for cold sweetening of potato tubers (MENÉNDEZ ET AL., 2002).

Table 3.2.1: Distribution of the associated SSCP fragments *invGE-6f* and *invGF-4d* present in the genotypes ‘Satina’, ‘Diana’, and ‘Theresa’.

Genotype	<i>invGE-6f</i>	<i>invGF-4d</i>
‘Satina’	1	1
‘Diana’	1	1
‘Theresa’	1	1

1=SSCP fragment is present.

Out of 80 cloned cDNA sequences per genotype, a sequence was defined as an allele when it was found twice in two independent PCRs⁴. The consensus sequence of all alleles found in one genotype was then used for allele definition when variable sequence polymorphisms occurred.

invGE and *invGF* invertase alleles obtained from each genotype are listed in Table 3.2.2.

Table 3.2.2: Overview of *invGE* and *invGF* alleles.

Genotype	Full-length <i>invGE</i> clones	<i>invGE</i> alleles	Full-length <i>invGF</i> clones	<i>invGF</i> alleles
‘Satina’	10	<i>E_SA</i> <i>E_SN1</i> <i>E_SN2</i> <i>E_SN3</i>	14	<i>F_SN1</i> <i>F_SN2</i> <i>F_SN3</i> <i>F_SN4</i>
‘Diana’	8	<i>E_DA</i> <i>E_DN1</i> <i>E_DN2</i>	4	<i>F_DN1</i> <i>F_DN2</i>
‘Theresa’	19	<i>E_TA</i> <i>E_TN1</i> <i>E_TN2</i> <i>E_TN3</i>	4	<i>F_TN1</i> <i>F_TN2</i>
P18	9	<i>E_P18N1</i> <i>E_P18N2</i>	2	<i>F_P18N</i>
P40	8	<i>E_P40N1</i> <i>E_P40N2</i>	4	<i>F_P40N1</i> <i>F_P40N2</i>
P54	5	<i>E_P54N1</i> <i>E_P54N2</i>	10	<i>F_P54N1</i> <i>F_P54N2</i>

The ‘A’ in the allele name stands for ‘association with better potato chips quality’ and refers to clones containing SNP 1103* (Table 3.2.3), which leads to a histidine at protein position 368. The ‘N’ in the allele name means ‘not associated’ with tuber starch and sugar content. The full-length clone number refers to the number of fully sequenced clones from each genotype used for allele definition. PCR amplification was carried out using gene specific full-length primers: *invGE*-*invGE*-F-fulgth/*invGE*-R-fulgth (chapter 2, Table 2.1.2); *invGF*-*invGF*-F-fulgth/*invGF*-R-fulgth (chapter 2, Table 2.1.2).

From all six genotypes analyzed in this study, 17 *invGE* and 13 *invGF* alleles were identified.

⁴ Exceptions are listed in Appendix A 3.2.

3.2.1.2.1 *invGE* cDNA alleles of the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’⁵

Cloning and sequencing of ‘Satina’ cDNA resulted in ten full-length clones, from which four *invGE* alleles *E_SA*, *E_SNI*, *E_SN2*, and *E_SN3* were defined. From the genotype ‘Diana’ eight full-length clones were obtained, and of those three different alleles *E_DA*, *E_DNI*, and *E_DN2* were identified. For the cultivar ‘Theresa’ 19 full-length clones were isolated and four alleles *E_TA*, *E_TN1*, *E_TN2*, and *E_TN3* were defined.

The nucleotide sequence comparison (Appendix A 3.2.13) of the alleles described above detected 125 SNPs, which resulted in 43 amino acid exchanges (Table 3.2.3; Figure 3.2.3).

Table 3.2.3: SNPs present in ‘Satina’, ‘Diana’, and ‘Theresa’ alleles

Position cDNA SNP	<i>E_SA</i>	<i>E_SNI</i>	<i>E_SN2</i>	<i>E_SN3</i>	<i>E_DA</i>	<i>E_DNI</i>	<i>E_DN2</i>	<i>E_TA</i>	<i>E_TN1</i>	<i>E_TN2</i>	<i>E_TN3</i>	aa
58	T	C	C	T	T	C	C	T	T	C	C	T/C F20L
85	A	G	G	A	A	G	G	A	G	G	G	A/G K29G
86	A	G	G	A	A	G	G	A	G	G	G	s.
95	G	C	C	C	G	C	C	C	C	C	C	G/C G32A
106	G	A	A	A	G	A	A	A	A	A	A	s.
108	T	T	A	T	T	T	A	T	T	T	T	A/T V36I
133	G	G	G	G	G	G	G	G	G	G	C	C/G P45A
135	T	T	T	A	T	T	T	A	T	T	T	s.
162	T	T	T	T	T	T	T	T	G	T	T	s.
163	G	G	G	A	G	G	G	A	G	G	G	A/G S55G
187	C	C	C	T	C	C	C	T	T	T	C	T/C Y63H
204	T	T	T	T	T	T	T	T	A	T	T	s.
231	A	A	A	A	A	A	A	A	G	A	A	s.
276	T	C	C	C	T	C	C	C	C	C	T	s.
345	C	T	T	T	C	T	T	T	T	T	T	s.
351	C	T	T	T	C	T	T	T	T	T	T	s.
390	C	A	A	A	C	A	A	A	A	A	A	s.
402	T	A	A	T	T	A	A	T	T	A	A	s.
411	G	A	A	A	G	G	A	A	A	G	A	s.
414	T	C	C	T	T	C	C	T	T	C	T	s.
415	G	G	A	G	G	G	A	G	G	G	G	A/G I139V
420	T	C	C	C	T	C	C	C	C	C	C	s.
429	T	C	C	C	T	C	C	C	C	C	C	s.

⁵ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: ‘Satina’ (Appendix A 3.2.1), ‘Diana’ (Appendix A 3.2.2), ‘Theresa’ (Appendix A 3.2.3). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *E_SA* (Appendix A 3.2.4), *E_SNI* (Appendix A 3.2.5), *E_SN3* (Appendix A 3.2.6), *E_DA* (Appendix A 3.2.7), *E_DNI* (Appendix A 3.2.8), *E_TA* (Appendix A 3.2.9), *E_TN1* (Appendix A 3.2.10), *E_TN2* (Appendix A 3.2.11), *E_TN3* (Appendix A 3.2.12). For the alleles *E_SN2* and *E_DN2* only one sequence was obtained, respectively. It has been shown that both alleles are real because allele specific SNPs were detected by pyrosequencing assay of cDNA and genomic DNA.

Position cDNA SNP	<i>E</i> <i>SA</i>	<i>E</i> <i>SN1</i>	<i>E</i> <i>SN2</i>	<i>E</i> <i>SN3</i>	<i>E</i> <i>DA</i>	<i>E</i> <i>DN1</i>	<i>E</i> <i>DN2</i>	<i>E</i> <i>TA</i>	<i>E</i> <i>TN1</i>	<i>E</i> <i>TN2</i>	<i>E</i> <i>TN3</i>	aa
444	T	C	C	C	T	C	C	C	C	C	C	s.
451	A	T	T	T	A	T	T	T	T	T	T	T/C T151S
456	A	A	A	G	A	A	A	G	G	A	A	s.
462	A	A	A	G	A	A	A	G	G	A	A	s.
465	C	T	T	T	C	T	T	T	T	T	T	s.
471	C	A	G	A	C	A	A	A	A	A	A	s.
477	A	G	G	G	A	G	G	G	A	G	G	s.
495	G	A	A	A	G	A	A	A	A	A	A	s.
511	G	A	A	A	G	A	A	A	A	A	A	G/A V171I
519	C	T	T	T	C	T	T	T	T	T	T	s.
531	G	G	A	G	G	G	A	G	G	G	G	s.
538	A	G	G	G	A	G	G	G	G	G	G	A/G I180V
540	T	T	A	A	T	A	A	A	T	A	T	s.
576	T	T	T	T	T	T	T	T	T	T	C	s.
588	T	A	A	T	T	A	A	T	T	A	A	s.
597	T	C	C	C	T	C	C	C	C	C	C	s.
598	G	C	C	C	G	C	C	C	C	C	C	s.
599	T	A	A	A	T	A	A	A	A	A	A	T/A V200Q
607	G	C	C	C	G	C	C	C	C	C	C	G/C V203L
615	G	A	A	A	G	A	A	A	A	A	A	s.
639	A	A	C	A	A	A	C	A	A	A	A	C/A N213K
645	A	A	A	A	A	A	A	A	T	A	A	T/A S215R
658	T	T	T	T	T	T	G	T	T	T	T	G/T V220L
688	G	G	A	G	G	G	A	G	G	G	G	A/G T230A/ V
689	T	C	C	C	T	C	C	C	C	C	C	T/C V230A/ T
702	C	T	T	T	C	T	T	T	T	T	T	s.
753	C	C	C	T	C	T	C	T	T	T	T	s.
768	T	A	A	A	T	A	A	A	A	A	A	T/A N256K
774	T	T	T	T	T	A	T	T	T	A	T	s.
780	T	C	C	C	T	C	C	C	C	C	C	s.
807	---	A	A	---	---	---	A	---	---	---	---	K269
808	---	A	A	---	---	---	A	---	---	---	---	K269
809	---	A	A	---	---	---	A	---	---	---	---	K269
810*	T	A	T	T	T	T	T	T	T	T	T	A/T K270N
817*	T	C	C	C	T	T	C	T	C	T	T	C/T H273Y
855*	T	G	T	G	T	G	G	T	T	G	G	T/G D285E
858*	C	T	T	T	C	T	T	C	T	T	T	s.
864*	T	T	T	T	T	T	A	T	T	T	T	s.
870*	C	T	T	T	C	T	T	C	T	T	T	s.
884*	G	A	A	A	G	A	A	G	A	A	A	G/A R295K

Position cDNA SNP	<i>E</i> <i>SA</i>	<i>E</i> <i>SN1</i>	<i>E</i> <i>SN2</i>	<i>E</i> <i>SN3</i>	<i>E</i> <i>DA</i>	<i>E</i> <i>DN1</i>	<i>E</i> <i>DN2</i>	<i>E</i> <i>TA</i>	<i>E</i> <i>TN1</i>	<i>E</i> <i>TN2</i>	<i>E</i> <i>TN3</i>	aa
909*	C	C	C	C	C	C	C	C	C	C	T	s.
924*	T	T	T	T	T	T	C	T	T	T	T	s.
926*	G	C	C	C	G	C	C	G	C	C	C	s.
927*	T	G	G	G	T	G	G	T	G	G	G	T/G C309S
945*	T	C	C	C	T	C	C	T	C	C	C	s.
975*	T	C	C	T	T	C	C	T	T	C	C	s.
981*	T	C	C	C	T	C	C	T	C	C	C	s.
983*	C	C	C	C	C	C	T	C	C	C	C	T/C L328P
986*	C	T	T	T	C	T	T	C	T	T	T	s.
987*	T	G	G	G	T	G	G	T	G	G	G	T/G T329M
1005*	G	A	A	A	G	A	A	G	A	A	A	s.
1010*	G	G	G	G	G	G	G	G	C	G	G	C/G A337G
1016*	C	C	C	C	C	C	C	C	T	C	C	T/C I340T
1017*	C	A	A	A	C	A	A	C	A	A	A	s.
1065*	T	T	T	A	T	T	T	T	A	T	T	s.
1068*	A	T	A	A	A	A	A	A	A	A	A	s.
1083*	G	G	G	A	G	G	G	G	A	G	G	s.
1086*	A	T	T	T	A	T	T	A	T	T	T	s.
1101*	T	C	C	C	T	C	C	T	C	C	C	s.
1103*	A	C	C	C	A	C	C	A	C	C	C	A/C H368P
1110*	T	T	T	C	T	T	T	T	T	T	T	s.
1117*	T	T	T	T	T	C	T	T	T	C	T	s.
1149*	C	C	A	A	C	A	A	C	A	A	A	s.
1152*	A	G	A	A	A	A	A	A	A	A	A	s.
1158*	A	G	G	G	A	G	G	A	G	G	G	s.
1167*	C	T	T	T	C	T	T	C	T	T	T	s.
1168*	---	---	A	---	---	A	A	---	---	A	---	I389
1169*	---	---	T	---	---	T	T	---	---	T	---	I389
1170*	---	---	T	---	---	T	T	---	---	T	---	I389
1179**	T	C	C	C	T	C	C	T	C	C	C	s.
1182**	C	C	C	C	C	C	C	C	C	C	G	G/C K394N
1191**	G	G	G	G	G	G	G	G	A	G	G	s.
1216**	A	A	A	G	A	A	A	A	G	A	A	G/A E406K
1237**	T	G	G	G	T	G	G	T	G	G	G	T/G S413A
1254**	A	G	G	G	A	G	G	A	G	G	G	s.
1257**	T	C	C	C	T	C	C	T	C	C	C	s.
1272**	G	G	A	G	G	G	A	G	G	G	G	s.
1276**	A	G	A	A	A	A	A	A	A	A	A	G/A E426K
1278**	A	G	G	G	A	G	G	A	G	G	G	s.
1281**	T	C	C	C	T	C	C	T	C	C	C	s.
1299**	T	A	A	A	T	A	A	T	A	A	A	T/A N433K
1305**	T	C	C	T	T	C	C	T	C	C	C	s.
1332**	C	C	C	C	C	C	T	C	C	C	C	s.
1335**	A	T	A	T	A	A	A	A	T	A	A	s.
1365**	A	G	A	A	A	A	G	A	A	A	A	s.

Position cDNA SNP	<i>E</i> <i>SA</i>	<i>E</i> <i>SN1</i>	<i>E</i> <i>SN2</i>	<i>E</i> <i>SN3</i>	<i>E</i> <i>DA</i>	<i>E</i> <i>DN1</i>	<i>E</i> <i>DN2</i>	<i>E</i> <i>TA</i>	<i>E</i> <i>TN1</i>	<i>E</i> <i>TN2</i>	<i>E</i> <i>TN3</i>	aa
1379**	C	C	T	C	C	T	C	C	C	T	T	T/C V460A
1380**	A	G	G	G	A	G	G	A	G	G	G	
1384**	T	T	T	G	T	T	T	T	T	T	T	G/T V462L
1395**	A	A	A	G	A	A	A	A	G	A	A	s.
1410**	T	C	C	C	T	C	C	T	C	C	C	s.
1413**	A	A	A	A	A	A	G	A	A	A	A	s.
1427**	G	G	G	G	G	G	G	G	G	G	A	A/G Q476R
1456**	G	A	G	G	G	G	G	G	G	G	G	A/G I486V
1476**	T	T	T	T	T	T	T	T	T	T	G	s.
1556**	T	T	C	T	T	C	C	T	T	C	T	C/T T519M
1617**	A	T	T	T	A	T	T	A	T	T	T	s.
1653**	A	G	G	G	A	G	G	A	G	G	G	s.
1659**	A	G	G	G	A	G	G	A	G	G	G	s.
1665**	T	T	T	T	T	C	C	T	T	T	C	s.
1666**	G	A	A	A	G	A	A	G	A	A	A	G/A D556N
1692**	T	C	T	T	T	T	T	T	T	T	T	s.
1723**	C	C	C	C	C	C	C	C	A	C	C	A/C I575L

SNP position numbering refers to cDNA sequence where ‘1’ represents the adenine of the start codon ATG and the polymorphism is described. * and **: SNP positions changed because of amino acid insertions, numbers refer to alignment nomenclature and not to the SNP positions of the cDNA for standardized comparison. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are coloured in orange.

The four ‘Satina’ alleles displayed a total of 95 SNPs, of which 30 caused an amino acid exchange. In the three allelic sequences of ‘Diana’ 87 SNPs occurred, from which 26 resulted in an amino acid exchange. The four ‘Theresa’ alleles differed in 70 SNPs. 22 of them led to an amino acid exchanges. The alleles *E_SN1*, *E_SN2*, and *E_DN2* contain the additional amino acid lysine (K) at position 259. An amino acid insertion also occurs at position 390* where the alleles *E_SN2*, *E_DN1*, *E_DN2*, and *E_TN2* contain an additional isoleucine (I).

The amino acid alignment shows the polymorphisms between all 11 *invGE* alleles from the three tetraploid potato cultivars. The comparison of these deduced protein sequences revealed 42 variable amino acid positions in the different genotypes, which are highlighted in colour (Figure 3.2.3).

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      *           20           *           40           *           60           *           80
E_DN1 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_TN2 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_TN3 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_SN2 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_DN2 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_SN1 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_SN3 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_TN1 : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_SA : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_DA : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86
E_TA : MELFMKSSSLWGLEIYLFCLFIVLSNINGVFAASHNIFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPMYNGVYHLFYQYNP : 86

      *           100           *           120           *           140           *           160           *
E_DN1 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_TN2 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_TN3 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_SN2 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_DN2 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_SN1 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_SN3 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_TN1 : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_SA : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_DA : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172
E_TA : KGSVWGNIVWAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSATILPNNKPVILYTGVDVSDHSQVQNYAIPANLSDPFLRKWIK : 172

      180           *           200           *           220           *           240           *           2
E_DN1 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_TN2 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_TN3 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_SN2 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_DN2 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_SN1 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_SN3 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_TN1 : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_SA : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_DA : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258
E_TA : PNNNPLIIPDINSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNT : 258

      60           *           280           *           300           *           320           *           340
E_DN1 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_TN2 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_TN3 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_SN2 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 344
E_DN2 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 344
E_SN1 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 344
E_SN3 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_TN1 : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_SA : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_DA : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343
E_TA : NGLDASYRGKLVKKVVLKNSLDVNRFFYYTIGMYDTTKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIVWGWTNESDV : 343

      *           360           *           380           *           400           *           420           *
E_DN1 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 429
E_TN2 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 429
E_TN3 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428
E_SN2 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 430
E_DN2 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 430
E_SN1 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 429
E_SN3 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428
E_TN1 : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428
E_SA : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428
E_DA : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428
E_TA : LPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEBELETLRKQKIQLNKKLSKGMFEVKGISASQADIEVSFSFSSLNKAQEF : 428

      440           *           460           *           480           *           500           *
E_DN1 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 515
E_TN2 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 515
E_TN3 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514
E_SN2 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 516
E_DN2 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 516
E_SN1 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 515
E_SN3 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514
E_TN1 : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514
E_SA : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514
E_DA : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514
E_TA : DPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLKMSDARRSTMRQNEAMYKPSFAGYVDVDL : 514

      520           *           540           *           560           *           580
E_DN1 : VDTKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 585
E_TN2 : VDTKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 585
E_TN3 : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
E_SN2 : VDTKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 586
E_DN2 : VDTKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 586
E_SN1 : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 585
E_SN3 : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
E_TN1 : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
E_SA : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
E_DA : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
E_TA : VDMKKLSLRSLLIDNSVVSFSGAGGKTCITSRVYPTLAIHNAHLFVFNNGSETITITETLNAWSMDVPKMH : 584
```

Figure 3.2.3: Amino acid alignment of ‘Satina’, ‘Diana’, and ‘Theresa’ *invGE* alleles. Amino acid exchanges are shown in colour. Additional amino acids are coloured in grey bars. At protein position 230 three different amino acids are displayed.

The amino acid alignment of *invGE* alleles of the tetraploid genotypes showed that ‘Satina’ and ‘Diana’ contain an allele identical at amino acid level ($E_SA=E_DA$).

3.2.1.2.2 *invGE* cDNA alleles of the diploid potato genotypes P18, P40, and P54⁶

From the genotype P18 nine full-length clones were obtained, and of those two alleles E_P18N1 and E_P18N2 were defined. Cloning and sequencing of P40 cDNA resulted in eight full-length clones, from which two different alleles E_P40N1 and E_P40N2 were identified. From the genotype P54 five full-length clones were isolated, and two alleles E_P54N1 and E_P54N2 were determined.

The alleles contain at nucleotide level (Appendix A 3.2.23) 48 SNPs. These sequence polymorphisms caused 23 amino acid exchanges (Table 3.2.4; Figure 3.2.4).

Table 3.2.4: SNPs present in P18, P40, and P54 alleles.

Position cDNA SNP	E_P18N1	E_P18N2	E_P40N1	E_P40N2	E_P54N1	E_P54N2	aa
58	C	C	C	T	C	T	T/C F20L
83	A	A	A	A	A	T	T/A I28N
85	G	G	G	A	G	A	A/G K29G
86	G	G	G	A	G	A	A/G K29G
108	A	T	A	T	T	T	s.
187	C	T	C	C	C	A	T/A/C Y63N/H
255	T	T	T	C	T	T	s.
402	A	A	A	T	A	T	s.
411	A	G	A	A	G	A	s.
414	C	C	C	T	C	T	s.
415	A	G	A	G	G	G	A/G I139V
426	C	C	C	T	C	C	s.
456	A	A	A	G	A	G	s.
462	A	A	A	G	A	G	s.
531	A	G	A	G	G	G	s.
540	A	A	A	T	A	T	s.
565	A	A	A	G	A	G	G/A E189K
619	G	G	G	A	G	G	A/G I207V
639	C	A	C	A	A	A	C/A N213K

⁶ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: P18 (Appendix A 3.2.14), P40 (Appendix A 3.2.15), P54 (Appendix A 3.2.16). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: E_P18N1 (Appendix A 3.2.17), E_P18N2 (Appendix A 3.2.18), E_P40N1 (Appendix A 3.2.19), E_P40N2 (Appendix A 3.2.20), E_P54N1 (Appendix A 3.2.21), E_P54N2 (Appendix A 3.2.22).

Position cDNA SNP	<i>E_P18N1</i>	<i>E_P18N2</i>	<i>E_P40N1</i>	<i>E_P40N2</i>	<i>E_P54N1</i>	<i>E_P54N2</i>	aa
645	A	A	A	A	A	G	s.
688	A	G	A	G	G	G	A/G T230A
753	C	T	C	T	T	C	s.
774	T	A	T	T	A	T	s.
807	A	---	A	---	---	---	K259
808	A	---	A	---	---	---	K259
809	A	---	A	---	---	---	K259
817*	C	T	C	C	T	C	T/C Y273H
855*	G	G	G	T	G	T	T/G D285E
975*	C	C	C	T	C	T	s.
1031*	T	T	T	T	T	G	G/T G344V
1101*	C	C	C	T	C	C	s.
1117*	T	C	T	T	C	T	s.
1168*	A	A	A	---	A	---	I389
1169*	T	T	T	---	T	---	I389
1170*	T	T	T	---	T	---	I389
1216**	A	A	A	G	A	G	G/A E406K
1272**	A	G	A	G	G	G	s.
1335**	A	A	A	T	A	T	s.
1379**	T	T	T	C	T	C	C/T A460V
1395**	A	A	A	G	A	G	s.
1462**	A	A	A	C	A	A	s.
1464**	G	G	G	A	G	G	A/G L488M
1471**	G	G	G	G	G	A	A/G N491D
1556**	C	C	C	T	C	T	T/C M519T
1616**	G	G	G	A	G	G	A/G D539G
1652**	C	C	C	C	C	T	T/C M551T
1665**	T	T	T	T	C	T	s.
1712**	C	C	C	T	C	C	T/C I561T

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. * and **: SNP positions changed because of amino acid insertions, numbers refer to alignment nomenclature and not to the SNP positions of the cDNA for standardized comparison. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are coloured in orange.

In the two allelic sequences of P18 12 SNPs occurred, from which five resulted in an amino acid exchange. The two P40 alleles differed in 30 SNPs. 14 of them led to changes in the protein sequence. From 26 SNPs of the two P54 alleles, 13 caused an amino acid substitution. The alleles *E_P18N1* and *E_P40N1* contain the additional amino acid lysine (K) at position 259. An amino acid insertion also occurs at position 390*, where the alleles *E_P18N1*, *E_P18N2*, *E_P40N1*, and *E_P54N1* contain an additional isoleucine (I).

The amino acid alignment displays the differences of the six cloned *invGE* alleles from the three diploid potato genotypes (Figure 3.2.4). The comparison of the protein sequences revealed 22 variable amino acid positions.

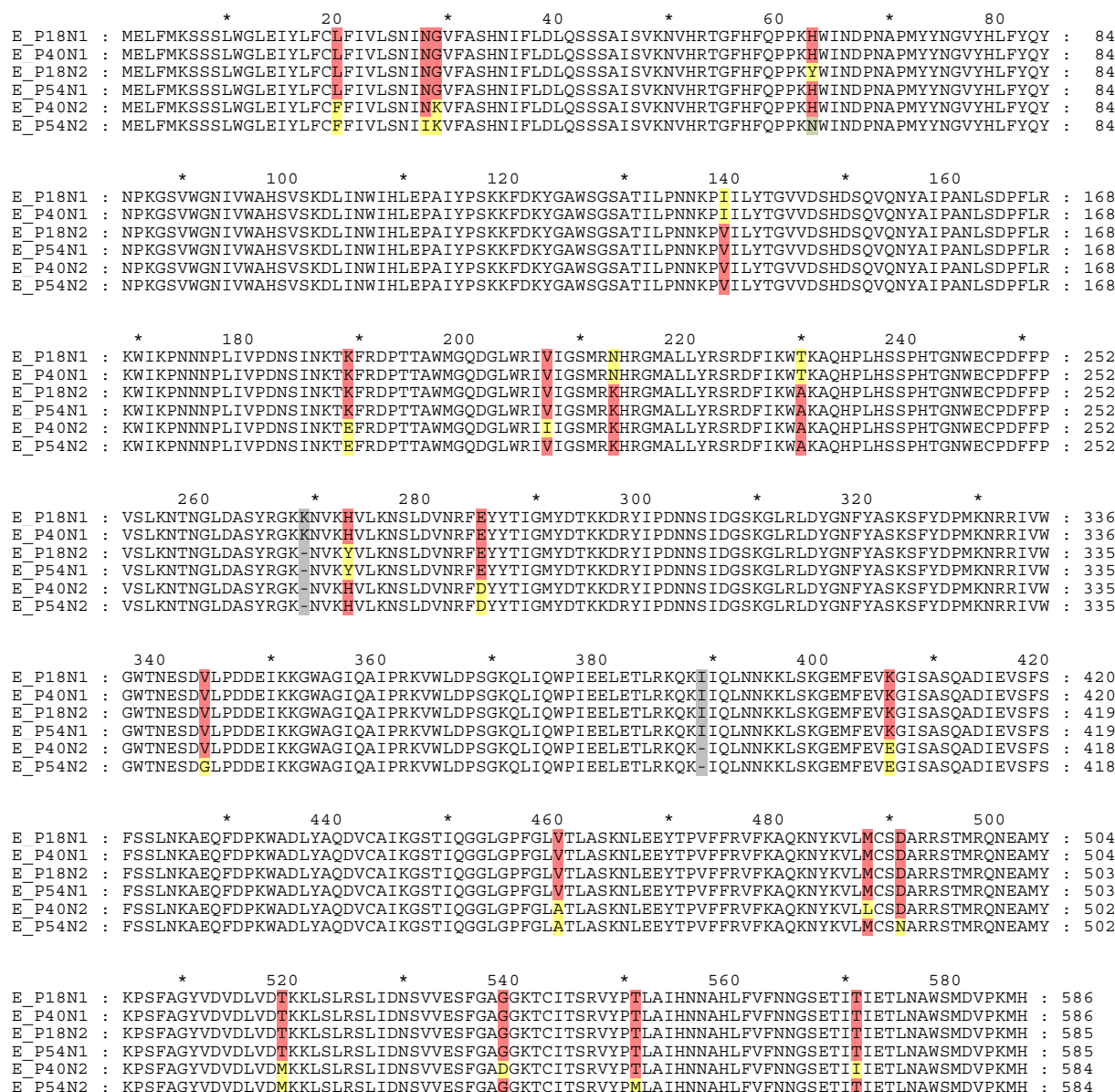


Figure 3.2.4: Amino acid alignment of P18, P40, and P54 *invGE* alleles. Amino acid exchanges are shown in colour. Additional amino acids are coloured in grey bars. At protein position 63 three different amino acids are displayed.

3.2.1.2.3 Amino acid alignment of all *invGE* invertase alleles of the analyzed potato genotypes

Multiple alignment of allelic *invGE* protein sequences revealed 53 variable amino acid polymorphisms between six genotypes (Figure 3.2.5). 21 amino acid exchanges were found to be genotype specific occurring only once.

[illegible]

Comparison of allelic cDNA sequences of the *invGE* gene from the three tetraploid and the three diploid genotypes revealed that different genotypes harbour amino acid sequence identical alleles. The alleles *E_SN2*, *E_P18N1*, and *E_P40N1*, the alleles *E_DN1* and *E_P54N1*, and the alleles *E_TN2* and *E_P18N2* have the same amino acid sequences.

In addition to the multiple amino acid alignment (3.2.1.2.3), the phenetic tree analysis was applied to group the invertase alleles according to their similarity at amino acid as well as at nucleotide level. Using the neighbour-joining method, the allelic classification visualized that *invGE* alleles from all six genotypes group in two clades and multiple subclades (Figure 3.2.6, Figure 3.2.7).

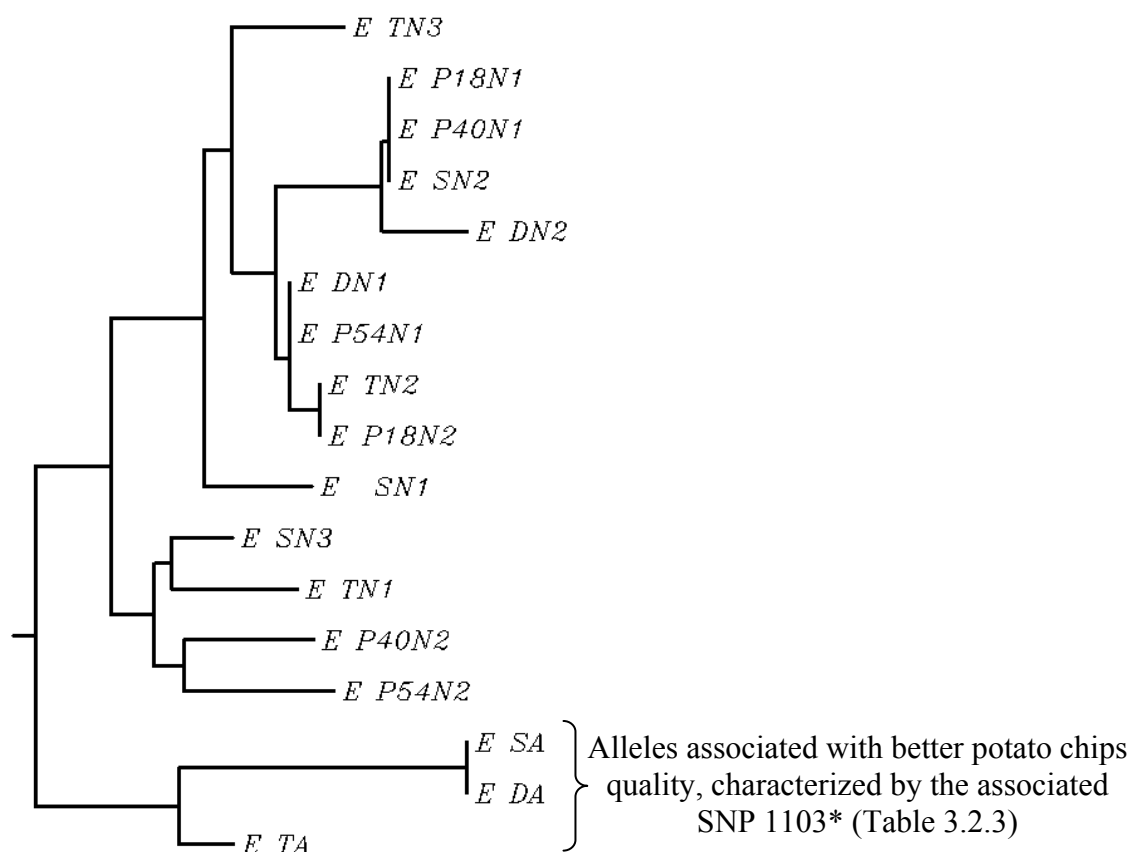


Figure 3.2.6: Amino acid based phenetic tree (Neighbour-joining method) of all cloned *invGE* invertase alleles.

The amino acid based phenetic tree has two clades. One clade contains most of the *invGE* alleles, and the other one the alleles *E_SA*, *E_DA*, and *E_TA*, which were found to be associated with superior potato chips quality.

The first clade then splits into diverse subclades due to the high number of amino acid polymorphisms between the different genotypes, although different genotypes contain amino acid sequence identical alleles. The alleles *E_SN2*, *E_P18N1*, and *E_P40N1*, the alleles *E_DN1* and *E_P54N1*, and the alleles *E_TN2* and *E_P18N2* have the same amino acid sequences.

The second clade consists of the alleles *E_SA*, *E_DA*, and *E_TA*. The two alleles *E_SA* and *E_DA* are identical at amino acid level, whilst *E_TA* differs in its amino acid sequence compared to the other two. However, all three alleles are characterized by the nucleotide adenine at cDNA SNP 1103* (Table 3.2.3), which is associated with better potato chips quality.

Cloning and sequencing of *invGE* alleles showed that different genotypes contain alleles identical at amino acid level but different regarding their nucleotide sequence. The allelic nucleotide composition was defined based on the consensus sequence of multiple alignments

of full-length clones obtained from each genotype (Table 3.2.2). Although SNPs are present at the mentioned cDNA positions, the nucleotide polymorphisms resulted in one and the same amino acid sequence.

Nucleotide comparison (Appendix A 3.2.24) showed that the associated alleles *E_SA* and *E_DA* consist of an identical nucleotide sequence as well as the alleles *E_TN2* and *E_P18N2*. The nucleotide sequences of the alleles and *E_P18N1* and *E_P40N1* are also identical, whilst the sequence of *E_SN2* differs at position 471. At position 471 a synonymous nucleotide exchange from A in the alleles *E_P18N1* and *E_P40N1* to G in allele *E_SN2* arose. Also one synonymous nucleotide variation was detected between the alleles *E_DN1* and *E_P54N1*. Allele *E_DN1* contains nucleotide A and allele *E_P54N1* consists of nucleotide G at position 1408**. Allele specific SNPs were detected in at least two sequences and used for comparing the allelic nucleotide polymorphisms.

The phenetic tree analysis showed the nucleotide polymorphisms between all alleles of the six different genotypes (Figure 3.2.7).

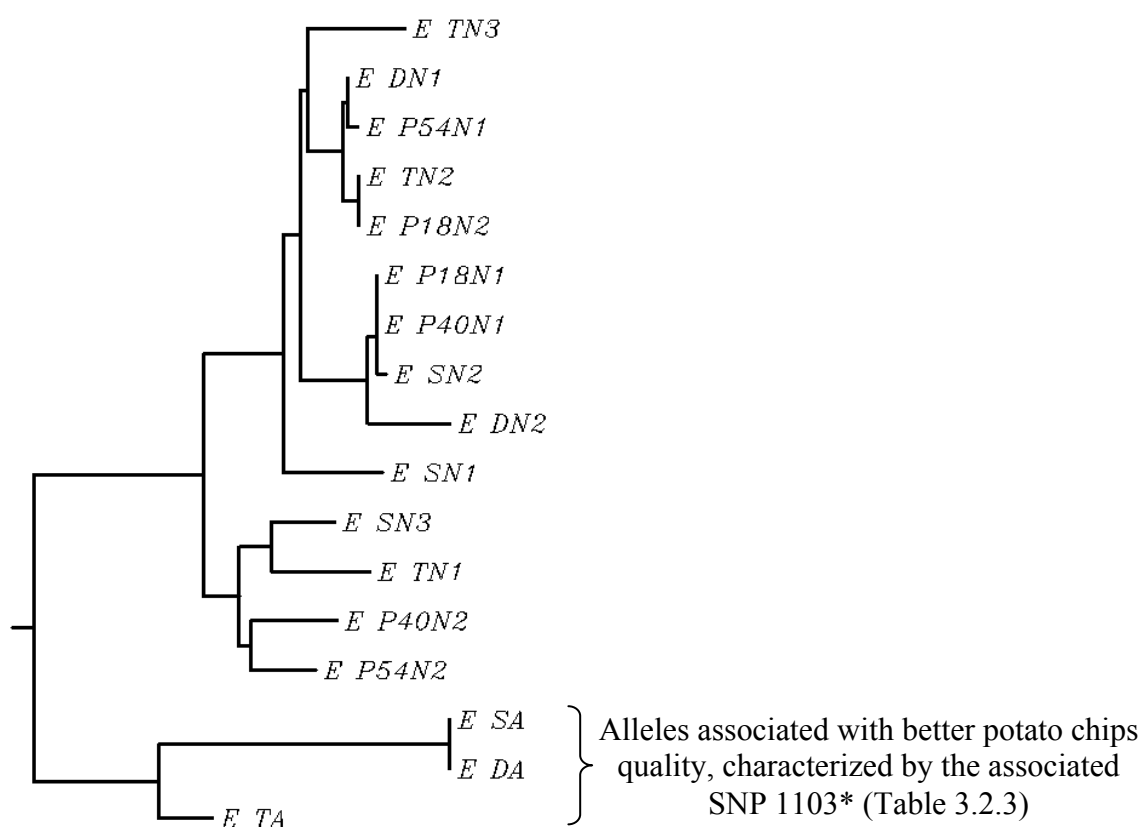


Figure 3.2.7: Nucleotide sequence based phenetic tree (Neighbour-joining tree) of all cloned *invGE* invertase alleles.

Although being very similar, the nucleotide based tree is characterized by an extended number of subclades due to a higher number of nucleotide polymorphisms in contrast to the amino acid based phenetic tree (Figure 3.2.6).

3.2.1.2.5 *invGF* cDNA alleles of the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’⁷

From the genotype ‘Satina’ 14 full-length clones were obtained, and of those four different alleles *F_SN1*, *F_SN2*, *F_SN3*, and *F_SN4* were identified. For ‘Diana’ four full-length clones were isolated, and two alleles *F_DN1* and *F_DN2* were determined. Cloning and sequencing of ‘Theresa’ cDNA resulted in four full-length clones, from which two *invGF* alleles *F_TN1* and *F_TN2* were defined.

The nucleotide comparison (Appendix A 3.2.35) of the alleles described above detected 86 SNPs, of which 19 were non synonymous and caused amino acid exchanges (Table 3.2.5; Figure 3.2.8).

Table 3.2.5: SNPs present in ‘Satina’, ‘Diana’, and ‘Theresa’ alleles.

Position cDNA SNP	<i>F_SN1</i>	<i>F_SN2</i>	<i>F_SN3</i>	<i>F_SN4</i>	<i>F_DN1</i>	<i>F_DN2</i>	<i>F_TN1</i>	<i>F_TN2</i>	aa
28	G	G	G	G	A	G	G	G	A/G T10A
58	A	A	A	G	A	A	G	A	G/A V20I
96	C	C	C	T	T	C	T	C	s.
111	T	C	C	C	C	C	C	C	s.
117	G	A	A	A	A	A	A	A	s.
141	A	C	A	C	C	A	C	A	A/C Q47H
162	C	C	C	T	T	C	T	C	s.
223	G	A	G	G	G	G	G	G	A/G I75V
225	C	T	T	C	C	C	C	C	s.
228	C	C	C	T	T	T	T	T	s.
249	C	C	C	C	C	T	C	T	s.
255	T	C	C	C	C	C	C	C	s.
273	T	C	T	C	C	T	C	T	s.
279	T	T	T	T	T	G	T	G	s.
345	C	T	T	T	T	C	T	C	s.
351	C	A	A	A	A	A	A	A	s.
363	C	C	C	T	T	C	T	C	s.
369	G	G	G	A	A	G	A	G	s.
378	A	A	G	A	A	A	A	A	s.

⁷ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: ‘Satina’ (Appendix A 3.2.25), ‘Diana’ (Appendix A 3.2.26), ‘Theresa’ (Appendix A 3.2.27). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *F_SN1* (Appendix A 3.2.28), *F_SN2* (Appendix A 3.2.29), *F_SN3* (Appendix A 3.2.30), *F_SN4* (Appendix A 3.2.31), *F_DN2* (Appendix A 3.2.32), *F_TN1* (Appendix A 3.2.33), *F_TN2* (Appendix A 3.2.34). For the allele *F_DN1* two nucleotide sequences were obtained, from which one showed an alternative stop codon leading to no termination of the sequence. Therefore, the nucleotide alignment of both sequences is not shown. *F_DN1* specific SNPs were shown to be real by pyrosequencing analysis of cDNA and genomic DNA.

Position cDNA SNP	<i>F_SN1</i>	<i>F_SN2</i>	<i>F_SN3</i>	<i>F_SN4</i>	<i>F_DN1</i>	<i>F_DN2</i>	<i>F_TN1</i>	<i>F_TN2</i>	aa
381	T	T	T	A	A	T	A	T	s.
405	G	G	G	A	A	G	A	G	s.
429	A	A	A	A	A	T	A	A	s.
432	A	A	A	G	G	A	G	A	s.
438	C	C	C	T	T	C	T	C	s.
459	C	T	C	C	C	C	C	C	s.
466	A	A	A	G	G	A	G	G	A/G I156V
478	T	G	G	A	A	T	A	T	T/G/A L160V/I
489	A	T	T	T	T	A	T	A	s.
491	T	T	T	T	T	T	T	C	C/T H164Y
495	C	C	C	A	A	T	A	T	s.
519	C	C	T	C	C	C	C	T	s.
542	C	A	A	C	C	C	C	C	A/C D181A
546	C	T	T	C	C	T	C	T	s.
555	G	A	G	G	G	G	G	G	s.
579	C	T	T	C	C	C	C	C	s.
582	A	A	A	T	T	A	T	A	s.
630	G	A	A	A	G	A	A	A	s.
651	T	A	A	T	T	T	T	T	s.
663	A	A	A	A	A	A	A	G	s.
672	T	T	T	C	C	T	C	T	s.
675	C	C	T	C	C	C	C	C	s.
717	T	T	T	C	C	T	C	C	s.
729	C	T	T	T	T	T	T	T	s.
750	C	T	T	C	C	C	C	C	s.
753	T	T	T	C	C	T	C	T	s.
778	A	A	A	C	C	A	C	A	s.
779	T	T	T	A	A	T	A	T	T/A I260Q
799	T	T	T	C	C	T	C	T	s.
801	T	T	T	C	C	T	C	T	C/T H267Y
805	T	T	T	T	T	T	T	A	A/T N269Y
820	A	A	A	A	A	A	A	G	G/A G274S
864	C	T	C	C	C	C	C	C	s.
876	G	A	A	A	A	A	A	A	s.
885	T	T	T	C	C	T	C	C	s.
886	G	G	G	A	A	G	A	G	A/G I296V
888	T	T	T	A	A	T	A	A	s.
903	T	T	C	T	T	T	T	T	s.
933	T	T	T	C	C	T	C	T	s.
958	A	A	A	T	T	A	T	A	T/A S320T
970	A	A	A	C	C	A	C	A	C/A P324T
981	C	C	C	T	T	C	T	C	s.
1029	G	A	A	G	G	G	G	G	s.
1038	T	T	T	C	C	T	C	T	s.
1041	G	G	G	T	T	G	T	G	s.
1047	A	A	A	A	A	A	A	G	s.

Position cDNA SNP	<i>F_SN1</i>	<i>F_SN2</i>	<i>F_SN3</i>	<i>F_SN4</i>	<i>F_DN1</i>	<i>F_DN2</i>	<i>F_TN1</i>	<i>F_TN2</i>	aa
1089	C	C	C	T	T	T	T	C	s.
1108	G	G	G	A	A	A	A	A	G/A V370I
1113	G	G	A	A	A	A	A	A	s.
1164	T	T	T	C	C	T	C	T	s.
1170	A	A	A	G	G	A	G	G	s.
1173	A	A	A	G	G	G	G	G	s.
1206	A	A	A	G	G	A	G	G	s.
1263	A	A	A	G	G	A	G	G	s.
1271	C	C	C	C	C	C	C	T	T/C L424S
1279	T	T	T	C	C	C	C	T	T/C S427P
1320	C	C	C	C	C	T	C	C	s.
1323	G	G	G	A	A	G	A	G	s.
1335	A	A	A	T	T	T	T	A	s.
1452	T	T	T	C	C	T	C	C	s.
1458	C	C	C	T	T	C	T	C	s.
1536	A	A	A	A	A	T	A	A	s.
1623	G	G	G	G	G	A	G	G	s.
1636	T	T	T	T	T	C	T	T	s.
1638	G	G	G	G	G	G	G	A	s.
1650	G	G	G	C	C	C	C	G	G/C E550D
1745	G	G	G	G	G	G	G	A	s.

SNP position numbering refers to cDNA sequence where ‘1’ represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are coloured in orange.

The four ‘Satina’ alleles differed in 70 SNPs, 14 of those led to amino acid exchanges. In the two allelic sequences of ‘Diana’ 47 SNPs were found, from which nine resulted in an amino acid substitution. The two alleles of ‘Theresa’ had a total of 47 SNPs, and 14 of those resulted in an amino acid difference.

The amino acid alignment shows the polymorphisms of all eight *invGF* alleles from the three tetraploid potato cultivars. The comparison of deduced protein sequences revealed 19 variable amino acid positions in the different genotypes displayed in coloured columns (Figure 3.2.8).

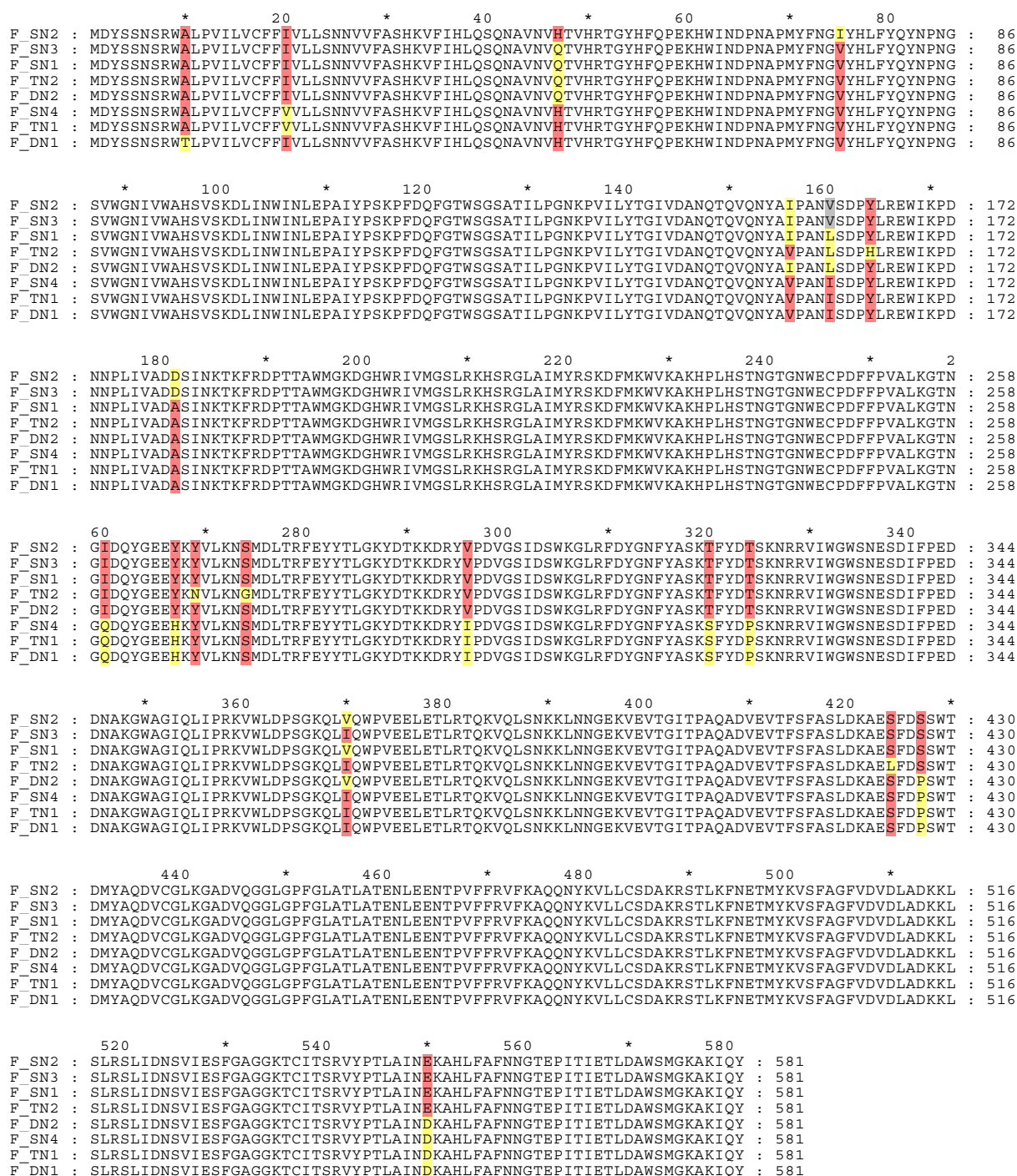


Figure 3.2.8: Amino acid alignment of ‘Satina’, ‘Diana’, and ‘Theresa’ *invGF* alleles. Amino acid exchanges are highlighted in colour. At protein position 160 three different amino acids are displayed.

Comparison of allelic cDNA sequences of the gene *invGF* from the three tetraploid cultivars revealed that the ‘Satina’ allele *F_SN4* and the ‘Theresa’ allele *F_TN1* are identical at amino acid level.

3.2.1.2.6 *invGF* cDNA alleles of the diploid potato genotypes P18, P40, and P54⁸

From P18 two full-length clones were isolated, and one allele *F_P18N* was determined. 20 partial sequenced P18 clones showed identical sequences to *F_P18N* but exhibited frame shifts or other modifications and, therefore, were not completely sequenced. From the genotype P40 four full-length clones were obtained, and of those two different alleles *F_P40N1* and *F_P40N2* were identified. Cloning and sequencing of P54 cDNA resulted in ten full-length clones, from which two *invGF* alleles *F_P54N1* and *F_P54N2* were defined.

31 SNPs for all alleles of the diploid genotypes were detected by nucleotide sequence comparison (Appendix A 3.2.43). Six of the identified polymorphisms caused an amino acid exchange (Table 3.2.6; Figure 3.2.9).

Table 3.2.6: SNPs present in P18, P40, and P54 alleles.

Position cDNA SNP	<i>F_P18N</i>	<i>F_P40N1</i>	<i>F_P40N2</i>	<i>F_P54N1</i>	<i>F_P54N2</i>	aa
111	T	T	C	T	T	s.
223	G	G	A	G	G	s.
225	C	C	T	C	C	T/C I75V
255	T	T	C	T	T	s.
351	C	C	A	C	C	s.
363	C	C	T	C	C	s.
542	C	C	A	C	C	A/C D181A
546	C	C	T	C	C	s.
579	C	C	T	C	C	s.
630	G	G	A	G	G	s.
651	T	T	A	T	T	s.
717	T	T	C	T	T	s.
729	C	C	T	C	C	s.
750	C	C	T	C	C	s.
791	G	G	A	G	G	A/G D264G
942	T	T	C	T	T	s.
1089	T	T	C	T	T	s.
1090	C	C	G	C	C	G/C A364P
1206	A	A	G	A	A	s.
1224	A	G	A	A	G	s.
1238	C	T	C	C	T	T/C I413T
1242	T	T	C	T	T	s.
1263	G	A	A	G	A	s.
1320	C	T	C	C	T	s.
1386	G	A	A	G	A	s.
1446	T	C	C	T	C	s.
1452	C	C	T	C	C	s.
1534	G	G	A	G	G	s.
1536	T	T	A	T	T	A/T T512A
1668	G	A	G	G	A	s.
1677	T	C	C	T	C	s.

⁸ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: P18 (Appendix A 3.2.36), P40 (Appendix A 3.2.37), P54 (Appendix A 3.2.38). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *F_P18N* (Appendix A 3.2.39), *F_P40N2* (Appendix A 3.2.40), *F_P54N1* (Appendix A 3.2.41), *F_P54N2* (Appendix A 3.2.42).

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are coloured in orange.

In the two allelic sequences of P40 27 SNPs occurred, six of those resulted in an amino acid exchange. The two P54 alleles differed in eight SNPs, and of those one led to an exchange in the amino acid sequence.

The amino acid alignment of the five *invGF* alleles from the three diploid potato genotypes showed the variability of six positions where amino acids differed (Figure 3.2.9).

F_P40N1	:	MDYSSNSRWALPVILVCF	20	FIVLLSNNVVF	40	FASHKVF	60	FIHLQSQNAVN	80	VQTVHRTGYHFQPEKHWINDPNAPMYFNG	:	84
F_P54N2	:	MDYSSNSRWALPVILVCF	20	FIVLLSNNVVF	40	FASHKVF	60	FIHLQSQNAVN	80	VQTVHRTGYHFQPEKHWINDPNAPMYFNG	:	84
F_P18N	:	MDYSSNSRWALPVILVCF	20	FIVLLSNNVVF	40	FASHKVF	60	FIHLQSQNAVN	80	VQTVHRTGYHFQPEKHWINDPNAPMYFNG	:	84
F_P54N1	:	MDYSSNSRWALPVILVCF	20	FIVLLSNNVVF	40	FASHKVF	60	FIHLQSQNAVN	80	VQTVHRTGYHFQPEKHWINDPNAPMYFNG	:	84
F_P40N2	:	MDYSSNSRWALPVILVCF	20	FIVLLSNNVVF	40	FASHKVF	60	FIHLQSQNAVN	80	VQTVHRTGYHFQPEKHWINDPNAPMYFNG	:	84
F_P40N1	:	NGSVWGNIVWAHSVSKDLINWINLEPAIYPSK	100	PFDPQFGTWSGSATILPGNKPVILYT	120	GTGIVDANQTQVQNYAIPANLSDPYLREW	:	168				
F_P54N2	:	NGSVWGNIVWAHSVSKDLINWINLEPAIYPSK	100	PFDPQFGTWSGSATILPGNKPVILYT	120	GTGIVDANQTQVQNYAIPANLSDPYLREW	:	168				
F_P18N	:	NGSVWGNIVWAHSVSKDLINWINLEPAIYPSK	100	PFDPQFGTWSGSATILPGNKPVILYT	120	GTGIVDANQTQVQNYAIPANLSDPYLREW	:	168				
F_P54N1	:	NGSVWGNIVWAHSVSKDLINWINLEPAIYPSK	100	PFDPQFGTWSGSATILPGNKPVILYT	120	GTGIVDANQTQVQNYAIPANLSDPYLREW	:	168				
F_P40N2	:	NGSVWGNIVWAHSVSKDLINWINLEPAIYPSK	100	PFDPQFGTWSGSATILPGNKPVILYT	120	GTGIVDANQTQVQNYAIPANLSDPYLREW	:	168				
F_P40N1	:	IKPDNNPLIVAD	180	ASINKTKFRDPTTAWMGKDGHWRIVMGS	200	SLRKHSRGLAIMYRSKDFMKVWKAKHPLHSTNGTGNWECDFFPV	:	252				
F_P54N2	:	IKPDNNPLIVAD	180	ASINKTKFRDPTTAWMGKDGHWRIVMGS	200	SLRKHSRGLAIMYRSKDFMKVWKAKHPLHSTNGTGNWECDFFPV	:	252				
F_P18N	:	IKPDNNPLIVAD	180	ASINKTKFRDPTTAWMGKDGHWRIVMGS	200	SLRKHSRGLAIMYRSKDFMKVWKAKHPLHSTNGTGNWECDFFPV	:	252				
F_P54N1	:	IKPDNNPLIVAD	180	ASINKTKFRDPTTAWMGKDGHWRIVMGS	200	SLRKHSRGLAIMYRSKDFMKVWKAKHPLHSTNGTGNWECDFFPV	:	252				
F_P40N2	:	IKPDNNPLIVAD	180	ASINKTKFRDPTTAWMGKDGHWRIVMGS	200	SLRKHSRGLAIMYRSKDFMKVWKAKHPLHSTNGTGNWECDFFPV	:	252				
F_P40N1	:	ALKGTNGIDQY	260	GEYKYVVLKNSMDLTRFEY	280	YTLGKYDTKKDRYVPDVGSIDSWKGLRFDYGNFYASKTFYDTSKNRRVIWGSN	:	336				
F_P54N2	:	ALKGTNGIDQY	260	GEYKYVVLKNSMDLTRFEY	280	YTLGKYDTKKDRYVPDVGSIDSWKGLRFDYGNFYASKTFYDTSKNRRVIWGSN	:	336				
F_P18N	:	ALKGTNGIDQY	260	GEYKYVVLKNSMDLTRFEY	280	YTLGKYDTKKDRYVPDVGSIDSWKGLRFDYGNFYASKTFYDTSKNRRVIWGSN	:	336				
F_P54N1	:	ALKGTNGIDQY	260	GEYKYVVLKNSMDLTRFEY	280	YTLGKYDTKKDRYVPDVGSIDSWKGLRFDYGNFYASKTFYDTSKNRRVIWGSN	:	336				
F_P40N2	:	ALKGTNGIDQY	260	GEYKYVVLKNSMDLTRFEY	280	YTLGKYDTKKDRYVPDVGSIDSWKGLRFDYGNFYASKTFYDTSKNRRVIWGSN	:	336				
F_P40N1	:	ESDIFPEDD	340	NAKGWAGIQLIPRKVWLD	360	PSGKQLVQWPVEELET	380	LRQKQVLSNKKLNNGEKVEVTGITPAQADVEV	400	YFSFASLD	:	420
F_P54N2	:	ESDIFPEDD	340	NAKGWAGIQLIPRKVWLD	360	PSGKQLVQWPVEELET	380	LRQKQVLSNKKLNNGEKVEVTGITPAQADVEV	400	YFSFASLD	:	420
F_P18N	:	ESDIFPEDD	340	NAKGWAGIQLIPRKVWLD	360	PSGKQLVQWPVEELET	380	LRQKQVLSNKKLNNGEKVEVTGITPAQADVEV	400	YFSFASLD	:	420
F_P54N1	:	ESDIFPEDD	340	NAKGWAGIQLIPRKVWLD	360	PSGKQLVQWPVEELET	380	LRQKQVLSNKKLNNGEKVEVTGITPAQADVEV	400	YFSFASLD	:	420
F_P40N2	:	ESDIFPEDD	340	NAKGWAGIQLIPRKVWLD	360	PSGKQLVQWPVEELET	380	LRQKQVLSNKKLNNGEKVEVTGITPAQADVEV	400	YFSFASLD	:	420
F_P40N1	:	KAESFDSSWTD	440	MYAQDVCG	460	LGKADVQGG	480	LGPPFGLATLATENLEENTPVFFRVFKAQQNYKVLLCSDAKRSTLKFNETMYKV	500	SFA	:	504
F_P54N2	:	KAESFDSSWTD	440	MYAQDVCG	460	LGKADVQGG	480	LGPPFGLATLATENLEENTPVFFRVFKAQQNYKVLLCSDAKRSTLKFNETMYKV	500	SFA	:	504
F_P18N	:	KAESFDSSWTD	440	MYAQDVCG	460	LGKADVQGG	480	LGPPFGLATLATENLEENTPVFFRVFKAQQNYKVLLCSDAKRSTLKFNETMYKV	500	SFA	:	504
F_P54N1	:	KAESFDSSWTD	440	MYAQDVCG	460	LGKADVQGG	480	LGPPFGLATLATENLEENTPVFFRVFKAQQNYKVLLCSDAKRSTLKFNETMYKV	500	SFA	:	504
F_P40N2	:	KAESFDSSWTD	440	MYAQDVCG	460	LGKADVQGG	480	LGPPFGLATLATENLEENTPVFFRVFKAQQNYKVLLCSDAKRSTLKFNETMYKV	500	SFA	:	504
F_P40N1	:	GFVDVDL	520	ADKKLSLRS	540	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPIT	560	ETLDAWSMGKAKIQY	580	:	581	
F_P54N2	:	GFVDVDL	520	ADKKLSLRS	540	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPIT	560	ETLDAWSMGKAKIQY	580	:	581	
F_P18N	:	GFVDVDL	520	ADKKLSLRS	540	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPIT	560	ETLDAWSMGKAKIQY	580	:	581	
F_P54N1	:	GFVDVDL	520	ADKKLSLRS	540	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPIT	560	ETLDAWSMGKAKIQY	580	:	581	
F_P40N2	:	GFVDVDL	520	ADKKLSLRS	540	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPIT	560	ETLDAWSMGKAKIQY	580	:	581	

Figure 3.2.9: Amino acid alignment of P18, P40, and P54 *invGF* alleles. Amino acid exchanges are highlighted in colour.

Comparison of allelic cDNA sequences of the gene *invGF* from the three diploid genotypes revealed that the alleles *F_P18N* and *F_P54N1*, and the alleles *F_P40N1* and *F_P54N2* are identical at amino acid level.

Figure 3.2.10: Amino acid alignment of all cloned *invGF* invertase alleles. Amino acid exchanges are highlighted in colour. At protein position 160 three different amino acids are displayed.

3.2.1.2.8 Phenetic trees of all *invGF* invertase alleles of the analyzed potato genotypes

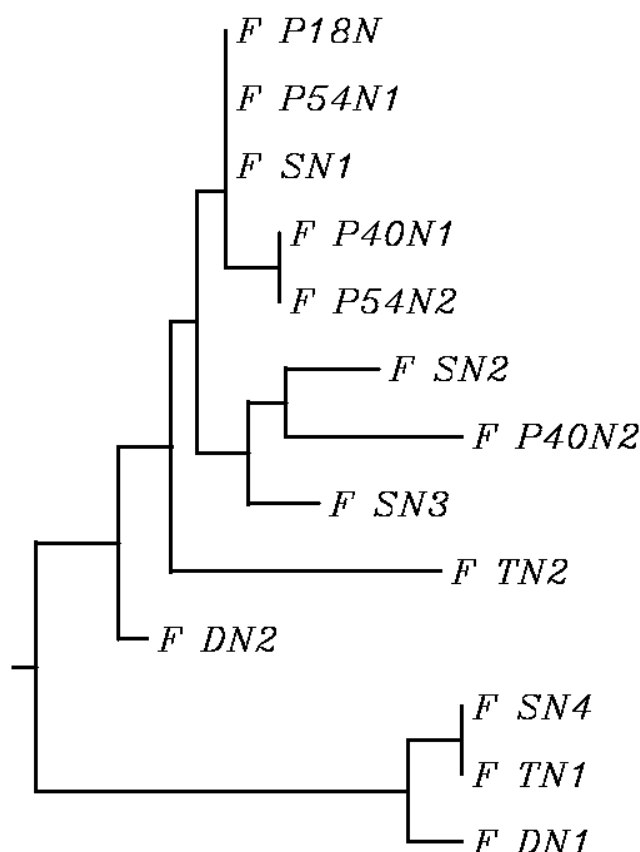


Figure 3.2.11: Amino acid based phenetic tree (Neighbour-joining method) of all cloned *invGF* invertase alleles.

The amino acid based phenetic tree splits into two main clades, from which the first one contains most of the *invGF* alleles. It is build of quite diverse subclades due to the high number of amino acid polymorphisms between the different genotypes. Amino acid comparison revealed sequence identical alleles in different genotypes. The alleles *F_SN1*, *F_P18N*, and *F_P54N1*, and the alleles *F_P40N1* and *F_P54N2* show the same amino acid sequence.

The second clade consists of the alleles *F_SN4*, *F_DN1*, and *F_TN1*. The two alleles *F_SN4* and *F_TN1* are identical at amino acid level, whilst *F_DN1* differs in its amino acid sequence. Cloning and sequencing of *invGF* alleles showed that different genotypes contain alleles identical at amino acid level but different at nucleic acid level (Appendix A 3.2.44). Table 3.2.7 shows a nucleotide comparison of amino acid identical alleles of different genotypes. The allelic nucleotide sequence was defined based on the consensus sequence of multiple alignments of full-length clones obtained from each genotype (Table 3.2.2). Although SNPs are present at the mentioned cDNA positions the nucleotide polymorphisms resulted in one and the same amino acid sequence.

Table 3.2.7: Genotype specific nucleotide differences of alleles *F_SNI*, *F_P18N*, and *F_P54N1*.

Position of cDNA SNP	<i>F_SNI</i>	<i>F_P18N</i>	<i>F_P54N1</i>
117	G	A	A
876	G	A	A
1089	C	T	T
1113	G	A	A
1173	A	G	G
1242	C	T	T
1263	A	G	G
1386	A	G	G
1446	C	T	T
1452	T	C	C
1536	A	T	T
1677	C	T	T

Nucleotide sequence comparison showed that the alleles *F_P40N1* and *F_P54N2*, and the alleles *F_SN4* and *F_TN1* were identical. The nucleotide sequences of the alleles *F_P18N* and *F_P54N1* were also identical, whilst the sequence of *F_SNI* differed in 12 positions. Allele specific SNPs were detected in at least two sequences and used for comparing the allelic nucleotide polymorphisms.

The nucleotide polymorphisms between all *invGF* alleles of the six analyzed genotypes were visualized using the phenetic tree analysis (Figure 3.2.12).

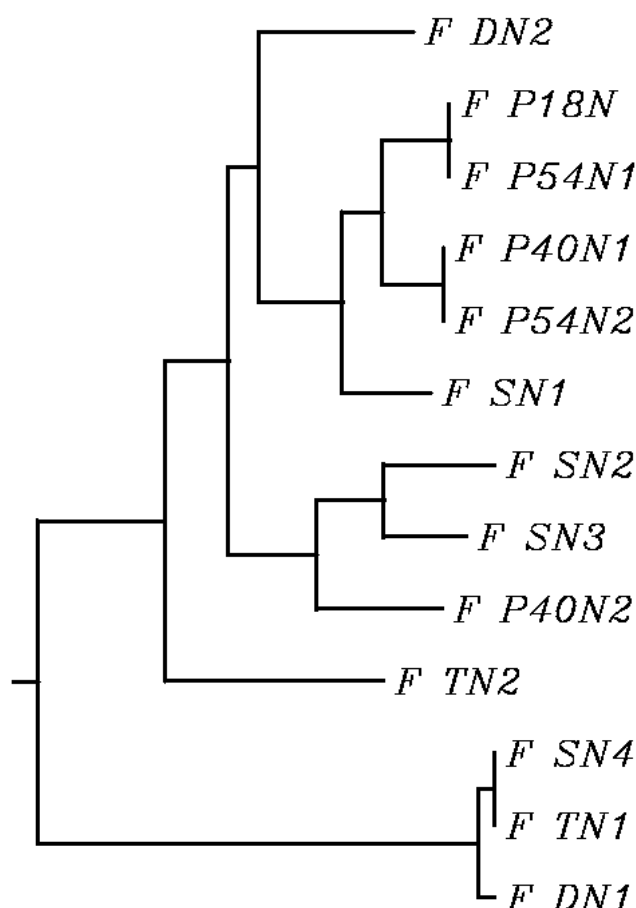


Figure 3.2.12: Nucleotide sequence based phenetic tree (Neighbour-joining tree) of all cloned *invGF* invertase alleles.

The alleles group regarding their similarity in the same clades and subclades as observed in the amino acid sequences based phenetic tree (Figure 3.2.11). Subclades are more subdivided because of nucleotide polymorphisms.

3.2.1.3 Three-dimensional modelling of *invGE* and *invGF* alleles

The 3D-modelling was performed by Pawel Durek, MPIMP/Golm. The allelic invertase molecular structures were modelled after the invertase 3D and crystal structure of cyanobacteria (ALBERTO ET AL., 2004). The models are comparative, superimposing two allelic sequences. Differences of the alleles are highlighted. The models include the putative sucrose binding site with the substrate sucrose. In addition to the structural visualization of amino acid exchanges, also the electrostatic potential (EP) of the molecules was mapped at pH 4.7 mimicking the apoplastic conditions.

❖ Structural modelling of *invGE* alleles of the cultivars ‘Satina’ and ‘Theresa’

Analysis of allelic molecular structures was performed with the associated ‘Satina’ allele *E_SA* compared to the allele *E_SN3*, and with the associated ‘Theresa’ allele *E_TA* compared to the allele *E_TN1*. The models contrast the allelic sequences with each other, meaning that

one sequence superimposes the other and vice versa (Figure 3.2.14). Based on protein sequence alignments (Figure 3.2.3), regions, which are affected directly or indirectly by amino acid exchanges, were identified (Figure 3.2.13).

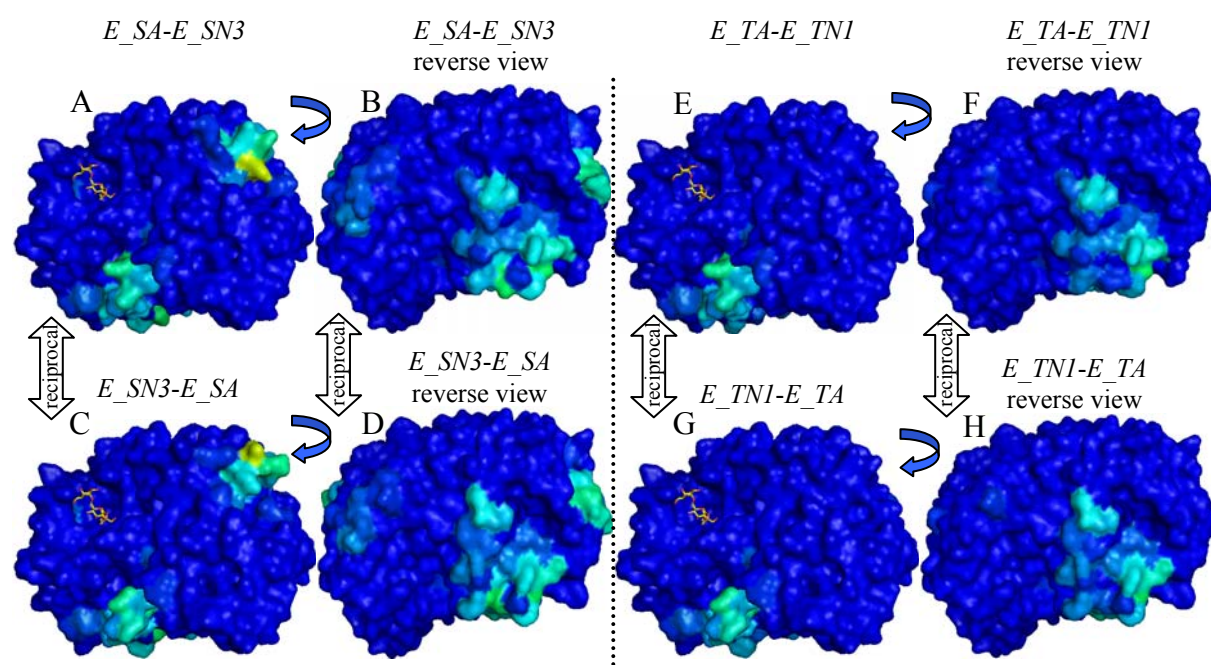
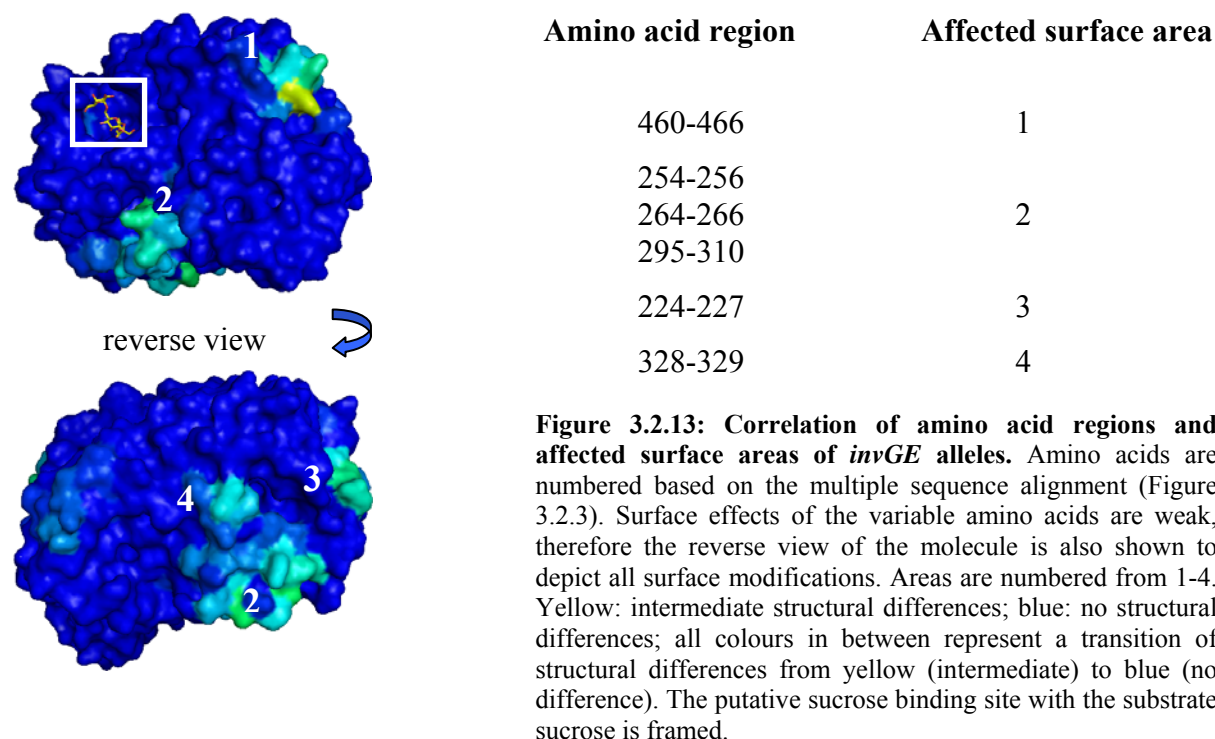


Figure 3.2.14: Structural comparison of ‘Satina’ alleles *E_SA*, *E_SN3* and ‘Theresa’ alleles *E_TA*, *E_TN1*. **A:** *E_SA* superimposes *E_SN3*; **B:** *E_SA* superimposes *E_SN3*, reverse view; **C:** *E_SN3* superimposes *E_SA*; **D:** *E_SN3* superimposes *E_SA*, reverse view; **E:** *E_TA* superimposes *E_TN1*; **F:** *E_TA* superimposes *E_TN1*, reverse view; **G:** *E_TN1* superimposes *E_TA*; **H:** *E_TN1* superimposes *E_TA*, reverse view. Yellow: intermediate structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from yellow (intermediate) to blue (no difference).

The models of the invertase alleles *E_SA*, *E_SN3* and *E_TA*, *E_TNI* showed structural differences on the enzyme's surface. The analysis of the allelic molecules *E_SA* and *E_SN3* revealed differences in regions 1, 2, 3, and 4 (Figure 3.2.13). Area 1 is defined by amino acids 460-466 showing alanine (A) in allele *E_SA* and valine (V) in allele *E_SN3* at position 462. Three amino acid exchanges between the two alleles occur in region 2 where at positions 256, 295 and 309 the protein sequence differs. The allele *E_SA* contains at position 256 asparagine (N) versus lysine (K) in allele *E_SN3*, at position 295 arginine (R) versus lysine (K), and at position 309 cysteine (C) versus serine (S). Region 3 is defined by amino acids 224-227. Here, the protein sequences of both 'Satina' alleles are identical. In the vicinity of this region, at position 230, an exchange from valine (V) to alanine (A) in the alleles *E_SA* and *E_SN3* respectively, occurs. This difference might influence surrounding areas and lead to variable protein folding. Amino acids 328 and 329 determine region 4, where at position 329 an amino acid exchange from threonine (T) in allele *E_SA* to methionine (M) in allele *E_SN3* appears. The models of the two 'Theresa' alleles *E_TA* and *E_TNI* show surface differences in the regions 2 and 4. Differences are absent in area 1 and 3 because of identical amino acid composition of both alleles. In region 2, which is defined by amino acids 254-256, 264-266, and 295-310 two amino acid exchanges occur. The allele *E_TA* contains at position 295 arginine (R) versus lysine (K) in allele *E_TNI*, and at position 309 cysteine (C) versus serine (S). These two substitutions are also present in allele *E_SA* from 'Satina'.

❖ Comparison of 3D-structures of the 'Satina' and 'Theresa' *invGE* alleles, which are associated with better chips quality

The tetraploid potato cultivars 'Satina' and 'Theresa' harbour alleles, which are not identical at amino acid level but exhibit the nucleotide adenine at cDNA SNP 1103* (Table 3.2.3), which is associated with better potato chips quality. Comparison of the 3D-structures of *Pain_SA* and *Pain_TA* molecules revealed minor surface differences between the two protein models (Figure 3.2.15).

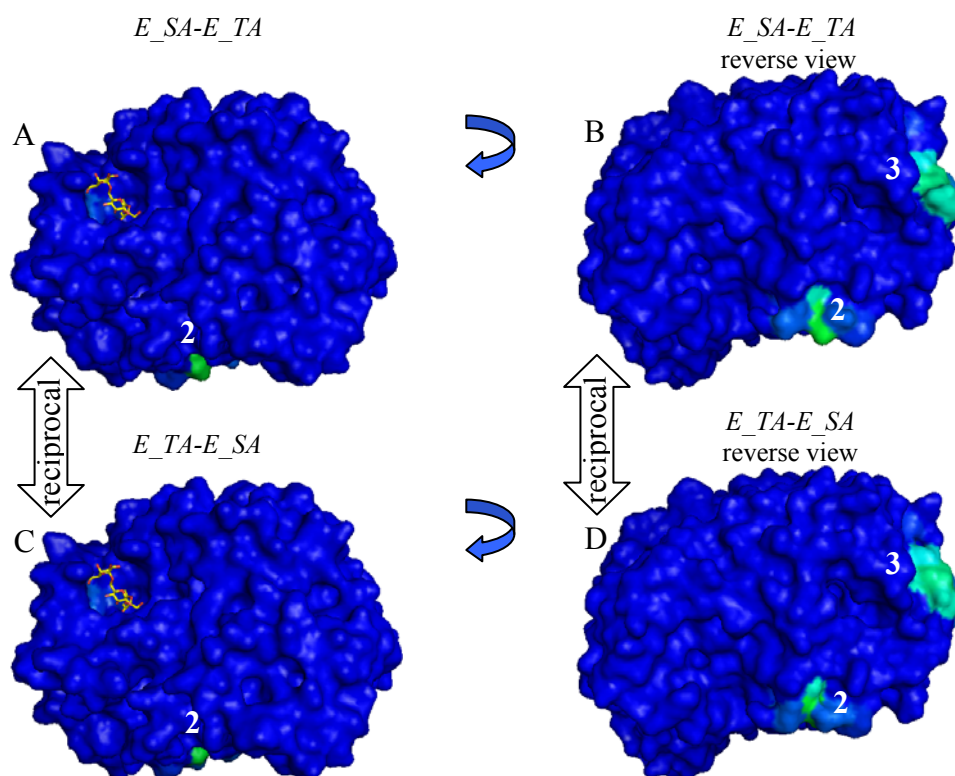


Figure 3.2.15: Comparative 3-D structures of the associated alleles *E_SA* and *E_TA*. A: *E_SA* superimposes *E_TA*; B: *E_SA* superimposes *E_TA*, reverse view; C: *E_TA* superimposes *E_SA*; D: *E_TA* superimposes *E_SA*, reverse view. Green: weak structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from green (weak) to blue (no difference).

Alleles *E_SA* and *E_TA* differ at 11 amino acid positions. The models of both allelic molecules revealed weak structural differences on the enzyme's surface in regions 2 and 3. Region 2 is defined by three amino acid stretches 254-256, where at position 256 an amino acid exchange was detected. The allele *E_SA* exhibits asparagine (N), whilst *E_TA* has lysine (K). Except this exchange, both alleles are identical in region 2 explaining the limited extension of this area compared to the other analyzed molecules (Figure 3.2.14). Region 3, defined by the amino acids 224-227, is identical in both alleles. The visible structural effect might be caused by the altered sequence at position 230 in the vicinity of region 3, which substitutes valine (V) to alanine (A) in the alleles *E_SA* and *E_TA*.

❖ Modelling the electrostatic potential (EP) of *invGE* alleles from the cultivars 'Satina' and 'Theresa'

The mapping of the EP of the *invGE* alleles *E_SA*, *E_SN3*, *E_TA*, and *E_TN1* revealed minor charge differences of the molecules at pH 4.7 (Figure 3.2.16).

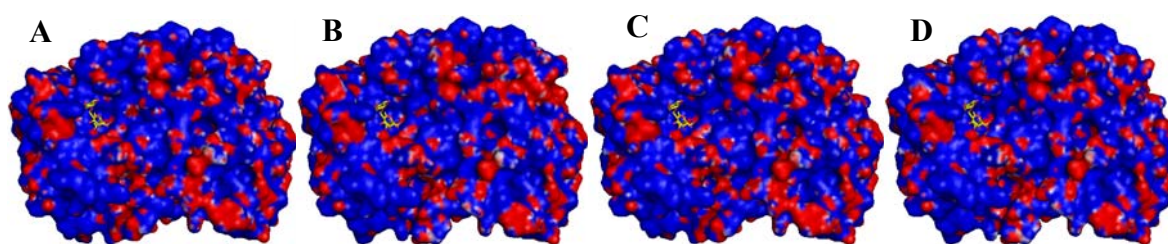


Figure 3.2.16: EP of the ‘Satina’ and ‘Theresa’ alleles. A: E_{SA} ; B: E_{SN3} ; C: E_{TA} ; D: E_{TN1} . Red: negatively charged; blue: positively charged; white: neutrally charged.

Focusing on the EP of the putative sucrose binding site

The putative sucrose binding site (Figure 3.2.13) is positively charged matching the partial negative charge of the substrate sucrose due to its hydroxyl groups.

Zooming into the putative sucrose binding domain revealed weak charge differences between the molecules E_{SA} (A) and E_{SN3} (B). The alleles E_{TA} (C) and E_{TN1} (D) showed no visible differences (Figure 3.2.17).

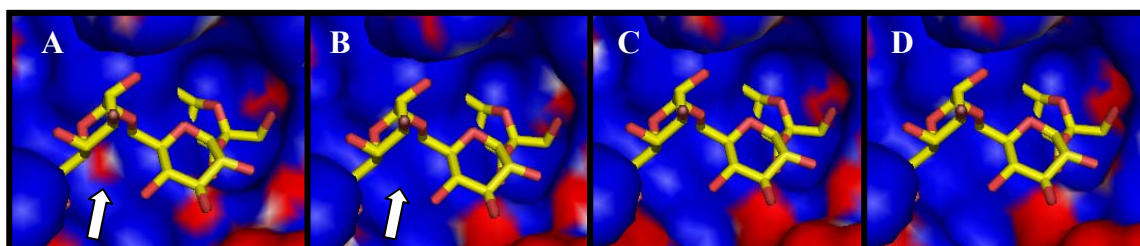


Figure 3.2.17: Focusing on the EP of the putative sucrose binding site of the ‘Satina’ and ‘Theresa’ alleles. A: E_{SA} ; B: E_{SN3} ; C: E_{TA} ; D: E_{TN1} . Red: negatively charged; blue: positively charged; white: neutrally charged.

The EP changes of the putative sucrose binding site of the two alleles from ‘Satina’ were weak. The EP of the associated allele E_{SA} (A) exhibits a small negatively charged area compared to the allele E_{SN3} (B). The charge of the ‘Theresa’ alleles E_{TA} (C) and E_{TN1} (D) were similar.

❖ Structural modelling of *invGF* alleles of the cultivar ‘Satina’

Molecular analysis of allelic structures was conducted using the two ‘Satina’ alleles F_{SN3} and F_{SN4} . The models compare the allelic sequences with each other by means of superimposing sequences of the different alleles (Figure 3.2.19). Doing so, differences between the alleles are visualized. Multiple alignment (Figure 3.2.8) of allelic protein sequences allowed to identify the regions, which are directly or indirectly affected by amino acid exchanges (Figure 3.2.18).

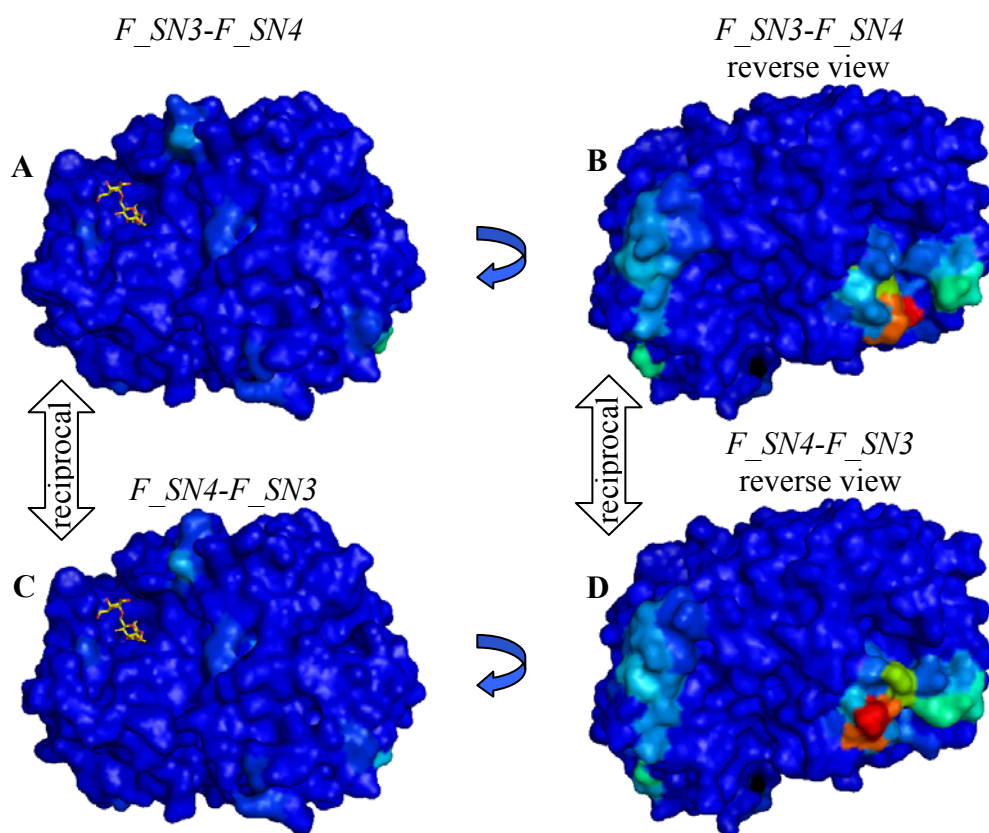
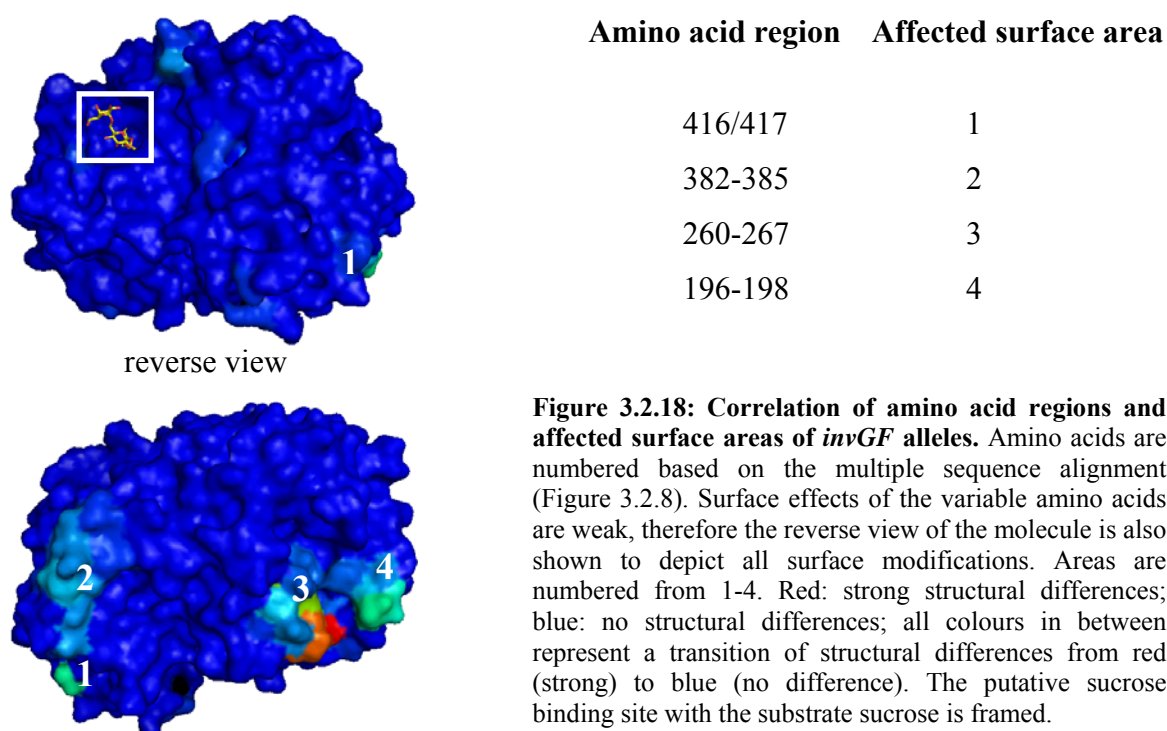


Figure 3.2.19: Structural comparison of ‘Satina’ alleles *F*_SN3 and *F*_SN4. **A:** *F*_SN3 superimposes *F*_SN4; **B:** *F*_SN3 superimposes *F*_SN4, reverse view; **C:** *F*_SN4 superimposes *F*_SN3; **D:** *F*_SN4 superimposes *F*_SN3, reverse view. Red: strong structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from red (strong) to blue (no difference).

The molecular models of the ‘Satina’ alleles *F_SN3* and *F_SN4* showed structural differences on the enzyme’s surface. Those differences were assigned to regions 1, 2, 3, and 4 (Figure 3.2.18). Area 1 is characterized by amino acids 416 and 417 exhibiting no differences between the two alleles. Region 2 is determined by the amino acids 382-385. Both alleles are identical in this region. Also area 4 consists of identical amino acids at positions 196-198. Although amino acids in the corresponding regions 1, 2, and 4 do not differ between the alleles, sterical differences occur. The reasons for the mild structural effects may be surrounding allelic sequences causing a variable lobing of the corresponding regions of the putative 3D invertase structure.

Region 3, which is defined by the amino acids 260-267 shows exchanges between the alleles. The allele *F_SN3* contains at position 260 isoleucine (I) versus glutamine (Q) in allele *F_SN4* and at position 267 tyrosine (Y) versus histidine (H).

❖ Modelling the electrostatic potential (EP) of *invGF* alleles from the cultivar ‘Satina’

The mapping of the EP of the *invGF* alleles *F_SN3* and *F_SN4* revealed minor charge differences of the molecules at pH 4.7 (Figure 3.2.20).

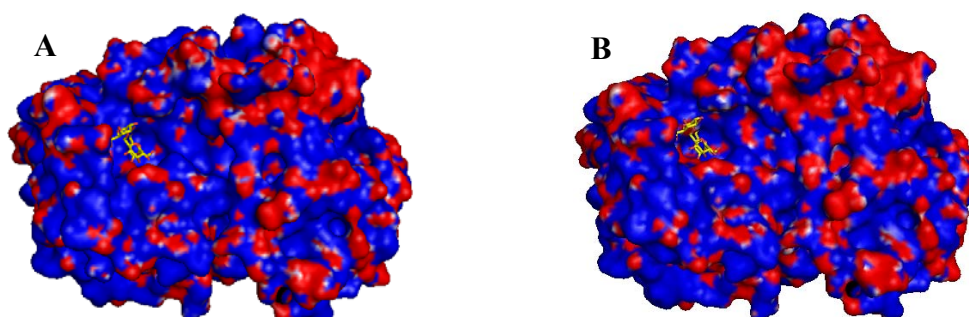


Figure 3.2.20: EP of the alleles *F_SN3* and *F_SN4*. **A:** *F_SN3*; **B:** *F_SN4*. Red: negatively charged; blue: positively charged; white: neutrally charged.

Focusing on the EP of the putative sucrose binding site

The putative sucrose binding site (Figure 3.2.18) has a positive charge matching the partial negative charge of the substrate sucrose due to its hydroxyl groups.

Zooming into the putative sucrose binding domain revealed charge differences between the molecules *F_SN3* (Figure 3.2.21, A) and *F_SN4* (Figure 3.2.21, B).

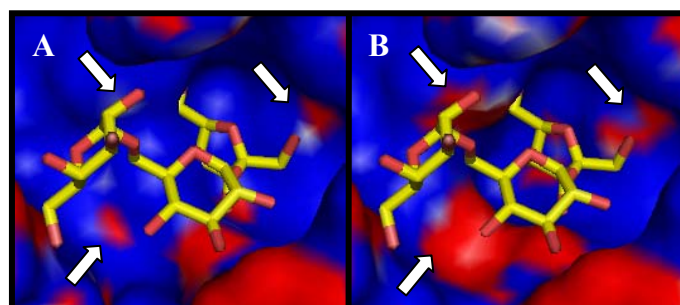


Figure 3.2.21: Focusing of the EP of the putative sucrose binding site of the alleles *F_SN3* and *F_SN4*. A: *F_SN3*; B: *F_SN4*. Red: negatively charged; blue: positively charged; white: neutrally charged.

The EP of the allele *F_SN4* (B) switched from a positive to a more negative charge caused by its allelic composition.

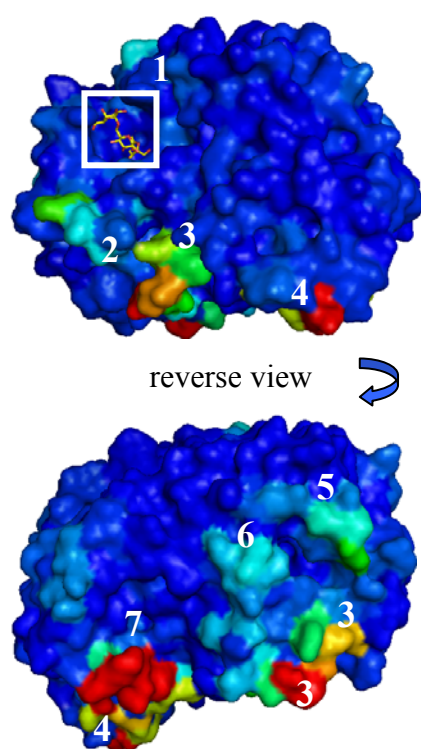
❖ Comparison of 3D-structures of *invGE* and *invGF* alleles

Association analysis showed that two SSCP fragments of the genes *invGE* (*invGE-6f*) and *invGF* (*invGF-4d*) are associated with better potato chips quality. These fragments are in LD showing nearly identical distributions within the population used (LI ET AL., 2005). Amino acid alignments (Figure 3.2.3) and phenetic tree analysis (Figure 3.2.6) showed the similarity of the three associated *invGE* alleles. Also three alleles of the gene *invGF* show high similarity to each other (Figure 3.2.8) and group in a separate clade compared to the other *invGF* alleles (Figure 3.2.11). Out of these three *invGF* alleles, which showed similar grouping as the associated *invGE* alleles and, therefore, might represent the associated *invGF* alleles, the allele *F_SN4* was selected for comparative 3D-modelling.

The associated *invGE* alleles *E_SA* and *E_TA* were compared against *F_SN4*, which was found to be amino acid and nucleotide identical to *F_TN1*. The *invGF* alleles *F_SN4*, *F_DN1*, and *F_TN1* are distinctive from the other *invGF* alleles.

Comparison of the 3D molecular structures of *E_SA* and *F_SN4*, and of *E_TA* and *F_SN4* revealed similar surface differences of both analyzed groups due to minor structural differences between the two associated *invGE* alleles *E_SA* and *E_TA*. The structural models are shown representatively from the comparison of *E_SA* and *F_SN4*.

A multiple alignment (Appendix A 3.2.45) of the allelic protein sequences allowed the identification of regions, which are directly or indirectly affected by amino acid exchanges (Figure 3.2.22).



Amino acid region	Affected surface area
86-89	1
133-135	2
210-213	
262-268	3
300-309	
419/420	4
476-480	
133-136	5
326-329	6
510-519	7

Figure 3.2.22: Correlation of amino acid regions and affected surface areas of comparative *invGE* and *invGF* alleles. Amino acids are numbered based on multiple sequence alignment (Appendix A 3.2.19). The reverse view of the molecule is shown because of the manifestation of allelic amino acid differences at the whole surface. Areas are numbered from 1-7. Red: strong structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from red (strong) to blue (no difference). The putative sucrose-binding site with the substrate sucrose is framed.

Comparison of the 3D molecular structures of *E_SA* and *F_SN4*, and of *E_TA* and *F_SN4* revealed surface differences between both analyzed enzymes (Figure 3.2.23).

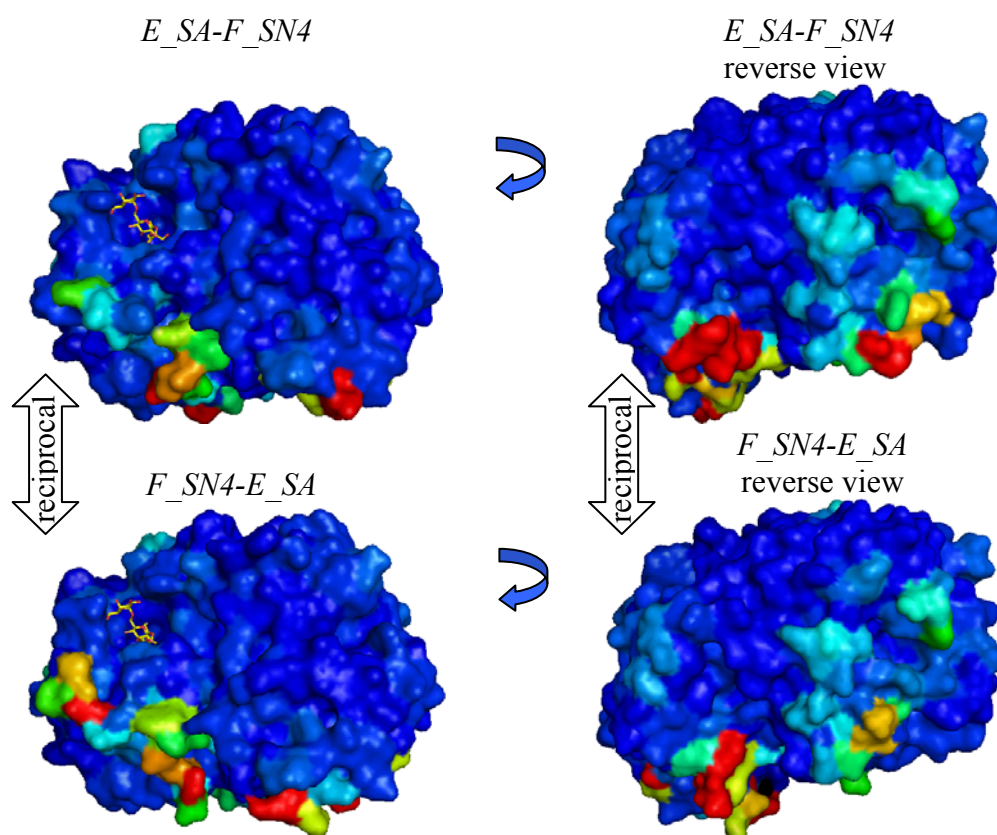


Figure 3.2.23: Structural comparison of *invGE* allele *E_SA* and *invGF* allele *F_SN4*. A: *E_SA* superimposes *F_SN4*; B: *E_SA* superimposes *F_SN4*, reverse view; C: *F_SN4* superimposes *E_SA*; D: *F_SN4* superimposes *E_SA*, reverse view. Red: strong structural differences; blue: no structural differences; all colours in between represent a transition of structural differences from red (strong) to blue (no difference).

The analysis of the corresponding allelic molecules revealed differences in the regions 1, 2, 3, 4, 5, 6, and 7 (area definition: Figure 3.2.22). Region 1 is defined by the amino acids 86-89. At position 87 an amino acid exchange occurs between both alleles. The allele *E_SA* contains lysine (K), whilst *F_SN4* consists of asparagine (N). The amino acids 133-135 and 210-213 represent the surface area 2. Amino acid exchanges at positions 135 and 211 cause structural differences. The allelic sequences change from asparagine (N135) in allele *E_SA* to glycine (G135) in allele *F_SN4*. At position 211 *E_SA* exhibits methionine (M) compared to leucine (L) in *F_SN4*. Region 3 is defined by the amino acids 262-268 and 300-309. Five regional amino acid exchanges are found between the two alleles. Protein sequence comparison revealed the following *E_SA* to *F_SN4* exchanges: A267Q, N302V, N303G, G307S and C308W. In the surface area 4 two amino acid substitutions occur. The first exchange is at position 420 where *E_SA* possesses serine (S), whilst *F_SN4* has alanine (A). The second exchange at position 480 shows a difference from lysine (K) in *E_SA* to glutamine (Q) in *F_SN4*. Region 5 is defined by the amino acids 133-136. At position 135 *E_SA* contains asparagine (N), *F_SN4* consists of lysine (G). The amino acids 326-329 are located in region 6 where an exchange at position 328 from threonine (T) in *E_SA* to serine (S) in *F_SN4* occurs. Region 7 is defined by the amino acids 510-519. In this area two exchanges at positions 515 and 517 were detected. The allele *E_SA* contains at position 515 valine (V) compared to alanine (A) in *F_SN4* and at position 517 a methionine (M), whilst *F_SN4* does not exhibit a comparable residue at this position.

The amino acid exchanges in the described regions manifest sterically and influence the lobing of the putative invertase molecules.

3.2.2 Functional characterization of the genes *invGE* and *invGF*

3.2.2.1 Differential expression analysis of *invGE* and *invGF* alleles

Referring to the expression pattern of the genes *invGE* and *invGF* observed by MADDISON ET AL. (1999), allele specific expression analysis was carried out in pyrosequencing assays using leaf tissues for *invGE* analysis and floral tissues for *invGF* characterization.

The pyrosequencing analysis was performed using cDNA from leaves and flowers and leaf and floral genomic DNA to determine the relative frequency of *invGE* and *invGF* alleles. Comparison of both samples revealed specific expression patterns not just for each genotype but also between different alleles of the same genotype. In the following Figures the relationship between the presence of alleles in the genome and their transcription levels are

illustrated. Accordingly, both values are shown together in terms of the relative expression level.

Additionally, plasmids harbouring one allele of complementary SNPs of corresponding genotypes were mixed in different ratios to monitor the accuracy of pyrosequencing analysis. Plasmid based measurements, working as positive controls, showed that SNP dependent variations of $\pm 5\%$ occurred. Values of cDNA and genomic dosages of the analysed alleles as determined by pyrosequencing were corrected for the observed SNP specific variations.

3.2.2.1.1 Expression patterns of *invGE* alleles in leaves of the tetraploid genotypes

By means of pyrosequencing the distribution of allele specific SNPs was analyzed (Table 3.2.3). The following Table 3.2.8 displays an overview of the SNPs selected from each tetraploid cultivar.

Table 3.2.8: Allele specific SNPs analyzed by pyrosequencing.

Genotype	Allele	SNP position	Allele specific SNP
'Satina'	<i>E_SA</i>	SNP 1237	<i>E_SA/E_SN1/E_SN2/E_SN3</i>
	<i>E_SN1</i>	SNP 1366	T/G/G/G
	<i>E_SN2</i>	SNP 1379	A/G/A/A
	<i>E_SN3</i>	SNP 1216	C/C/T/C
			A/A/A/G
'Diana'	<i>E_DA</i>	SNP 1086	<i>E_DA/E_DN1/E_DN2</i>
	<i>E_DN1</i>	SNP 1117	A/T/T
	<i>E_DN2</i>	SNP 924	T/C/T
			T/T/C
'Theresa'	<i>E_TA</i>	SNP 1615	<i>E_TA/E_TN1/E_TN2/E_TN3</i>
	<i>E_TN1</i>	SNP 1720	A/T/T/T
	<i>E_TN2</i>	SNP 1553	C/A/C/C
	<i>E_TN3</i>	SNP 1473	T/T/C/T
			T/T/T/G

SNP positions refer to cDNA sequence where '1' represents the adenine of the start codon ATG. Primers used for pyrosequencing are listed in chapter 2 (Table 2.1.5 and Table 2.2.9). Allele specific SNPs are highlighted in bold capitals.

The pyrosequencing analysis was performed using cDNA from leaves and leaf genomic DNA to measure the dosages of the alleles (Figure 3.2.24).

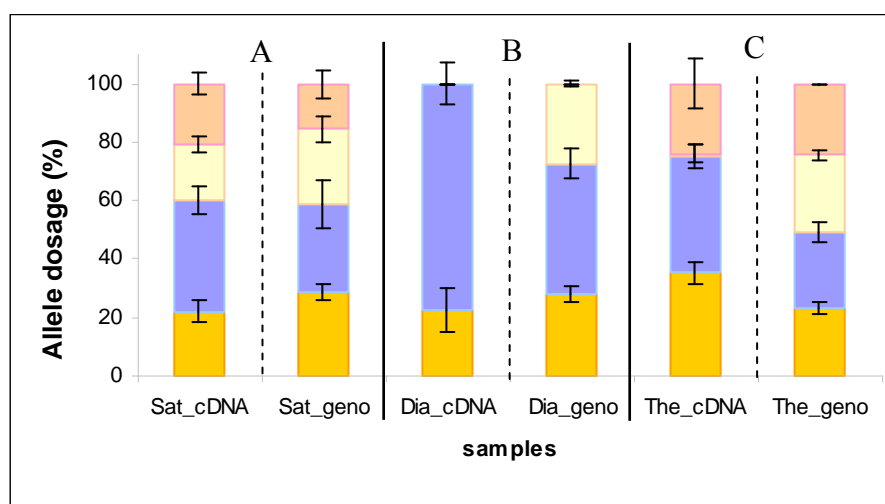


Figure 3.2.24: Pyrosequencing analysis of *invGE* alleles from the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’. **A:** Pyrosequencing of ‘Satina’ alleles. ■ : *E_SA*; ■ : *E_SN1*; ■ : *E_SN2*; ■ : *E_SN4*. **B:** Pyrosequencing of ‘Diana’ alleles. ■ : *E_DA*; ■ : *E_DN1*; ■ : *E_DN2*. **C:** Pyrosequencing of ‘Theresa’ alleles. ■ : *E_TA*; ■ : *E_TN1*; ■ : *E_TN2*; ■ : *E_TN3*. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of leaves. Percentages of the expression level of the cDNA samples are related to the genomic dosage of the alleles. Standard deviations are derived from two biological replicates done in technical triplicates.

Pyrosequencing analysis of genomic DNA showed that the three associated alleles *E_SA*, *E_DA*, and *E_TA* from the different cultivars are present in simplex (25%) in the corresponding genotype. For the cultivar ‘Satina’ four *invGE* alleles were found. If the alleles were transcribed according to the allele dosage, each allele is expected to contribute 25% of the total transcripts. Comparison of genomic allele dosage and allele transcription revealed that the alleles *E_SN1* and *E_SN4* were slightly over-represented, whilst the alleles *E_SA* and *E_SN2* were less abundant in cDNA samples.

In the cultivar ‘Diana’ three alleles were identified. Pyrosequencing analysis of genomic ‘Diana’ DNA showed allele dosages of 25% *E_DA*, 25% *E_DN2* and 50% of *E_DN1*.

However, the fraction represented by *E_DN1* could consist of two alleles since ‘Diana’ is tetraploid. In the pyrosequencing assay of ‘Diana’ leaf cDNA no transcripts for the allele *E_DN2* were detected due to a transcription rate below detection level, whilst the cDNA fraction represented by *E_DN1* showed a strong over-representation up to 80%. The abundance of the associated allele *E_DA* decreased from 25% of its genomic dosage to approximately 20% in cDNA.

The cultivar ‘Theresa’ has four *invGE* alleles *E_TA*, *E_TN1*, *E_TN2*, and *E_TN3*. Each allele is present in simplex (25%). Pyrosequencing analysis of leaf cDNA showed that the transcripts of the allele *E_TN3* were below the detection level. The abundance of the associated allele *E_TA* increased slightly from 25% genomic dosage to about 38% in leaf

cDNA. The strongest uprating was observed for the allele *E_TN1* that increased up to 40% when compared to the genomic dosage. Expression of the allele *E_TN2* did not differ compared to its genomic dosage and remained at 25%.

3.2.2.1.2 Expression pattern of *invGE* alleles in leaves of the diploid genotypes

For allele specific separation within the diploid potato genotypes P18, P40, and P54 allele specific SNPs were identified based on multiple nucleotide alignments (Table 3.2.4). SNP at position 108 was used to separate the two P18 alleles *E_P18N1* and *E_P18N2*. Using the SNP at position 58 the P40 alleles *E_P40N1*, *E_P40N2* and the P54 alleles *E_P54N1*, *E_P54N2* were distinguished (Table 3.2.9).

Table 3.2.9: Allele specific SNPs analyzed by pyrosequencing.

Genotype	Allele	SNP position	Allele specific SNP
P18	<i>E_P18N1</i>	SNP 108	A
	<i>E_P18N2</i>		T
P40	<i>E_P40N1</i>	SNP 58	C
	<i>E_P40N2</i>		T
P54	<i>E_P54N1</i>	SNP 58	C
	<i>E_P54N2</i>		T

SNP positions refer to cDNA sequence where '1' represents the adenine of the start codon ATG. Primers used for pyrosequencing are listed in chapter 2 (Table 2.1.5 and Table 2.2.10).

Pyrosequencing analysis was performed using cDNA from leaves (Figure 3.2.25).

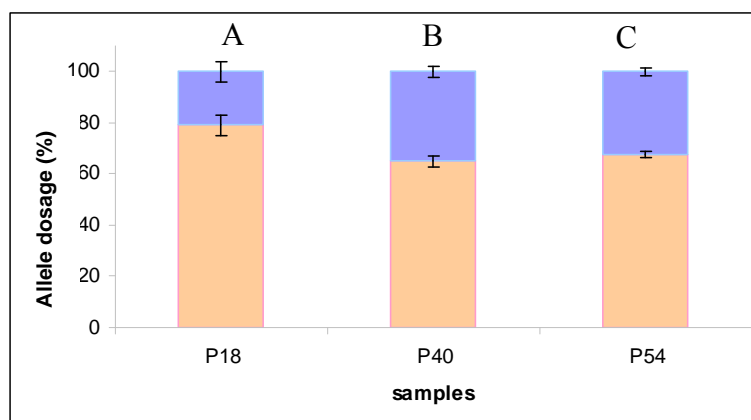


Figure 3.2.25: Pyrosequencing analysis of *invGE* alleles from the diploid genotypes P18, P40, and P54. A: Pyrosequencing of P18 alleles. ■: *E_P18N1*; ■: *E_P18N2*. **B:** Pyrosequencing of P40 alleles. ■: *E_P40N1*; ■: *E_P40N2*. **C:** Pyrosequencing of P54 alleles. ■: *E_P54N1*; ■: *E_P54N2*. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. Percentages of the expression level of the cDNA samples are related to the genomic allelic distribution being 50% because of the heterozygous diploid genotype. Genomic DNA of the corresponding genotypes was not analyzed in pyrosequencing assay. Standard deviations are derived from two biological replicates done in technical triplicates.

The three potato genotypes are diploid, meaning that they can be homozygous harbouring one *invGE* allele or heterozygous consisting of two *invGE* alleles. For P18, P40, and P54 two different *invGE* alleles were identified, respectively. Consequently, each allele per genotype is

present at 50%. Pyrosequencing analysis of leaf cDNA showed transcriptional changes of the genotype specific *invGE* alleles compared to their allele dosage. The allele *E_P18N1* revealed an increased abundance from 50% of its genomic distribution to 80% in cDNA.

In P40 the allele *E_P40N1* was under-represented in cDNA samples compared to its genomic dosage, whilst the allele *E_P40N2* showed an abundance increase up to 62% compared to its genomic dosage.

The two P54 alleles exhibit a similar expression pattern to P40. The allele order was inverted meaning that *E_P54N1* was over-represented, whilst *E_P54N2* was under-represented compared to the genomic allele dosage.

3.2.2.1.3 Expression pattern of *invGF* alleles in flowers of the tetraploid genotypes

Allele specific SNPs were selected based on multiple nucleotide alignment to separate different *invGF* alleles in the tetraploid genotypes (Table 3.2.5). The following Table 3.2.10 summarizes the selected SNPs from each cultivar.

Table 3.2.10: Allele specific SNPs analyzed by pyrosequencing.

Genotype	Allele	SNP position	Allele specific SNP
			<i>F_SN1/F_SN2/F_SN3/F_SN4</i>
‘Satina’	<i>F_SN1</i>	SNP 111	T /C/C/C
	<i>F_SN2</i>	SNP 459	C/ T /C/C
	<i>F_SN3</i>	SNP 378	A/A/ G /A
	<i>F_SN4</i>	SNP 96	C/C/C/ T
‘Diana’	<i>F_DN1</i>	SNP 96	T
	<i>F_DN2</i>		C
‘Theresa’	<i>F_TN1</i>	SNP 96	T
	<i>F_TN2</i>		C

SNP positions refer to cDNA sequence where ‘1’ represents the adenine of the start codon ATG. Primers used for pyrosequencing are listed in chapter 2 (Table 2.1.6 and Table 2.2.11). Allele specific SNPs are highlighted in bold capitals.

The pyrosequencing analysis was performed using cDNA and genomic DNA from flowers to measure the dosages of the alleles (Figure 3.2.26).

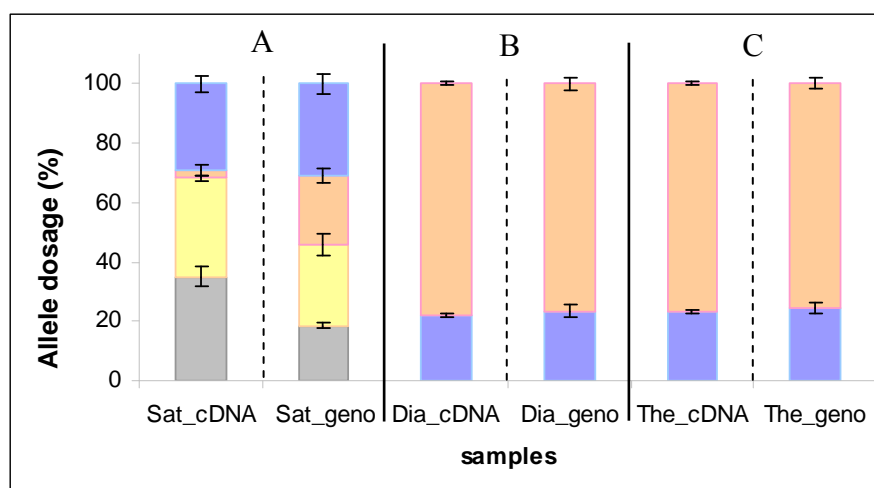


Figure 3.2.26: Pyrosequencing analysis of *invGF* alleles from the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’. **A:** Pyrosequencing of ‘Satina’ alleles. \square : *F_SN1*; \square : *F_SN2*; \square : *F_SN3*; \square : *F_SN4*. **B:** Pyrosequencing of ‘Diana’ alleles. \square : *F_DN1*; \square : *F_DN2*. **C:** Pyrosequencing of ‘Theresa’ alleles. \square : *F_TN1*; \square : *F_TN2*. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of flowers. Percentages of the expression level of the cDNA samples are related to the genomic allelic distribution. Standard deviations are derived from two biological replicates done in technical triplicates.

For the cultivar ‘Satina’ four *invGF* alleles were found. If the alleles were transcribed according to the allele dosage, each allele is expected to contribute 25% of the total transcripts. In the cDNA pyrosequencing assay marginal transcript was detectable for the allele *F_SN3*. The alleles *F_SN1* and *F_SN2* were more abundant in cDNA compared to their genomic dosage, whilst the presence of allele *F_SN4* remained stable at 25% according to its genomic distribution.

In the cultivar ‘Diana’ two alleles were identified. The allele *F_DN1* was present in simplex (25%), whilst the allele *F_DN2* was triplex (75%). However, *F_DN2* could possibly consist of up to three diverse alleles due to the tetraploidy of ‘Diana’. Pyrosequencing of flower cDNA revealed no differences in the expression pattern of both alleles compared to their genomic dosage.

The alleles *F_TN1* and *F_TN2* were defined for the cultivar ‘Theresa’. The allele *F_TN1* existed in simplex (25%), whilst the allele *F_TN2* was present in triplex (75%). Considering the tetraploidy, the fraction represented by the allele *F_TN2* might consist of more than one allele. Similar to the expression pattern of the two ‘Diana’ alleles, also the ‘Theresa’ alleles did not show any changes in their expression compared to their genomic dosage.

3.2.2.1.4 Expression pattern of *invGF* alleles in flowers of the diploid genotypes

From the potato genotype P18 one *invGF* allele was identified, whilst for P40 and P54 two alleles were found. In the pyrosequencing assay the distribution of allele specific SNPs were

analyzed (Table 3.2.6). In the following Table 3.2.11 discriminative SNPs of P40 and P54 alleles are listed.

Table 3.2.11: Allele specific SNPs analyzed by pyrosequencing.

Genotype	Allele	SNP position	Allele specific SNP
P40	<i>F_P40N1</i>	SNP 1534	G
	<i>F_P40N2</i>		A
P54	<i>F_P54N1</i>	SNP 1446	T
	<i>F_P54N2</i>		C

SNP positions refer to cDNA sequence where 1 represents the adenine of the start codon ATG. Primers used for pyrosequencing are listed in chapter 2 (Table 2.1.6 and Table 2.2.12).

The pyrosequencing analysis was performed using cDNA and genomic DNA from flowers to measure the dosages of the alleles (Figure 3.2.27)

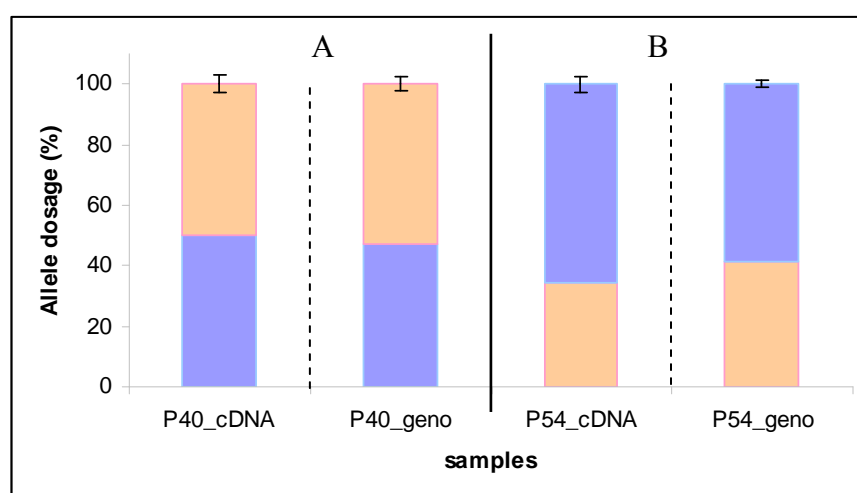


Figure 3.2.27: Pyrosequencing analysis of *invGF* alleles from the diploid genotypes P40 and P54. A: Pyrosequencing of P40 alleles. ■: *F_P40N1*; ■: *F_P40N2*. **B:** Pyrosequencing of P54 alleles. ■: *F_P54N1*; ■: *F_P54N2*. 100% is defined as the sum of alternative peaks measuring allele specific SNPs. In addition to pyrosequencing analysis of cDNA samples, the genomic dosage of the alleles was determined using genomic DNA of flowers. Percentages of the expression level of the cDNA samples are related to the genomic allelic distribution. Standard deviations are derived from two biological replicates done in technical triplicates.

Regarding the diploidy of the potato genotypes, the two alleles per genotype are expected to be equally frequent at genomic level. The expression pattern of the two P40 alleles *F_P40N1* and *F_P40N2* remained unchanged compared to their genomic dosage. Transcript levels of both alleles were consistent at approximately 50%.

In P54 the allele *F_P54N1* showed a decreased abundance of 15% compared to its genomic dosage. The allele *F_P54N2* was over-represented up to approximately 65% in cDNA compared to its genomic dosage.

3.2.2.2 Functional complementation of the yeast invertase mutant *SUC2*

Similar to the previous functional complementation analysis of the yeast invertase mutant *SUC2* with *Pain-1* cDNA alleles (section 3.1.2.2), cDNA alleles of the genes *invGE* and *invGF* were used for yeast transformation. Transforming *SUC2* with *invGE* and *invGF* cDNA alleles resulted in yeast transformants that were able to grow on sucrose as carbohydrate source, indicating functional complementation of the *SUC2* mutation.

The *invGE* and *invGF* cDNA alleles listed below (Figure 3.2.28) were used for complementation of *SUC2* and subsequent analysis of invertase activity (section 3.2.2.3).

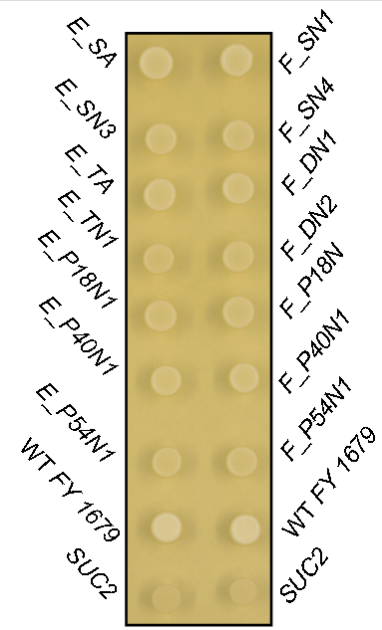
Genotype	<i>invGE</i> alleles	
'Satina'	<i>E_SA</i>	
	<i>E_SN1</i>	
	<i>E_SN3</i>	
'Theresa'	<i>E_TA</i>	
	<i>E_TN1</i>	
P18	<i>E_P18N1</i>	
P40	<i>E_P40N1</i>	
	<i>E_P40N2</i>	
P54	<i>E_P54N1</i>	
	<i>E_P54N2</i>	
'Satina'	<i>invGF</i> alleles	
	<i>F_SN1</i>	
	<i>F_SN2</i>	
	<i>F_SN3</i>	
'Diana'	<i>F_SN4</i>	
	<i>F_DN1</i>	
	<i>F_DN2</i>	
P18	<i>E_P18N</i>	
P40	<i>F_P40N1</i>	
	<i>F_P40N2</i>	
P54	<i>F_P54N1</i>	
	<i>F_P54N2</i>	
		<p>Representative yeast transformants complemented with <i>invGE</i> and <i>invGF</i> invertase alleles were spotted on solid yeast minimal broth with 2% sucrose as carbon source. The wild type <i>FY 1479</i> was plated as positive control, whilst the invertase mutant <i>SUC2</i> was the negative control.</p>

Figure 3.2.28: cDNA alleles used for *SUC2* complementation and *SUC2* transformants on solid yeast minimal media.

All *SUC2* invertase transformants exhibited substantial growth on sucrose and were used for the biochemical characterization of the *invGE* and *invGF* alleles. The potato alleles complemented the invertase deficiency of the yeast mutant not this good as the wild type *FY 1479* and displayed less colonies.

3.2.2.3 Biochemical characterization of *invGE* and *invGF* alleles

Putative 3D-models of allelic *invGE* and *invGF* molecules (section 3.2.1.3) indicated structural and electrostatic differences between the alleles, which could cause functional

differences. To test, whether structural characteristics might influence enzymatic activity of *invGE* and *invGF* invertase alleles, biochemical characterization was performed.

From yeast *SUC2* transformants complemented with *invGE* and *invGF* cDNA alleles, total protein was extracted. To test invertase activity, a modified protocol based on ZRENNER ET AL. (1995) was used. Invertase assays were carried out at 30°C. The *invGE* and *invGF* alleles displayed in Table 3.2.12 were used for biochemical characterization.

Table 3.2.12: *invGE* and *invGF* alleles used for biochemical characterization.

Genotype	<i>invGE</i> alleles	Genotype	<i>invGF</i> alleles
‘Satina’	<i>E_SA</i>	‘Satina’	<i>F_SN1</i>
	<i>E_SN1</i>		<i>F_SN2</i>
	<i>E_SN3</i>		<i>F_SN3</i>
‘Theresa’	<i>E_TA</i>	‘Diana’	<i>F_DN1</i>
	<i>E_TN1</i>		<i>F_DN2</i>
P18	<i>E_P18N1</i>	P18	<i>F_P18N</i>
P40	<i>E_P40N1</i>	P40	<i>F_P40N1</i>
	<i>E_P40N2</i>		<i>F_P40N2</i>
P54	<i>E_P54N1</i>	P54	<i>F_P54N1</i>
	<i>E_P54N2</i>		<i>F_P54N2</i>

The biochemical parameters Michaelis constant (K_m) and the maximal velocity (v_{max}) of invertase reaction were determined.

K_m values are independent from the enzyme concentration, whereas v_{max} values depend on the enzyme concentration. Due to the lack of an appropriate antibody against *invGE* and *invGF* cell wall-bound invertase isoforms, immunoblot analysis could not be performed (section 3.2.2.4). Therefore, v_{max} values of *invGE* and *invGF* alleles cannot be taken into account until protein quantification can be carried out.

❖ Biochemical characterization of *invGE* alleles

Results of the biochemical analysis for *invGE* alleles are shown in Table 3.2.13.

Table 3.2.13: K_m (mM) of *invGE* invertase alleles.

Genotype	<i>invGE</i> allele	K_m
‘Satina’	<i>E_SA</i>	19.66±0.93
	<i>E_SN1</i>	16.99±0.85
	<i>E_SN3</i>	23.97±0.93
‘Theresa’	<i>E_TA</i>	21.32±0.93
	<i>E_TN1</i>	18.64±0.85
P18	<i>E_P18N1</i>	24.4±0.93
P40	<i>E_P40N1</i>	17.13±0.85
	<i>E_P40N2</i>	17.1±0.85
P54	<i>E_P54N1</i>	22.73±0.85
	<i>E_P54N2</i>	19.51±0.85
<i>FY 1679</i>		24.25±2.5

Standard deviations are derived from three biological replicates for the associated alleles *E_SA* and *E_TA*, and the wild type reference strain *FY 1679* and from two biological replicates for the other alleles done in technical replicates to obtain six measurements. To make assays of different invertase isoforms comparable, the yeast reference strain *FY 1679* was used as positive control.

Table 3.2.14 summarizes the significance values for differences between the K_m values measured for the *invGE* alleles. The statistical calculation was performed by Benjamin Stich, MPIZ/Köln.

Table 3.2.14: Statistical significance levels of K_m values from the *invGE* invertase alleles.

Allele	<i>E_SA</i>	<i>E_SNI</i>	<i>E_SN3</i>	<i>E_TA</i>	<i>E_TNI</i>	<i>E_P18NI</i>	<i>E_P40NI</i>	<i>E_P40N2</i>	<i>E_P54NI</i>	<i>E_P54N2</i>
<i>E_SA</i>	---	0.056	0.006	0.23	0.43	0.0036	0.067	0.065	0.032	0.91
<i>E_SNI</i>	---	---	0.0001	0.0050	0.19	7.66E-05	0.92	0.93	0.0004	0.06
<i>E_SN3</i>	---	---	---	0.067	0.001	0.75	0.0002	0.0001	0.34	0.004
<i>E_TA</i>	---	---	---	---	0.055	0.038	0.006	0.006	0.29	0.18
<i>E_TNI</i>	---	---	---	---	---	0.0006	0.23	0.22	0.005	0.48
<i>E_P18NI</i>	---	---	---	---	---	---	8.97E-05	8.70E-05	0.21	0.002
<i>E_P40NI</i>	---	---	---	---	---	---	---	0.98	0.0005	0.07
<i>E_P40N2</i>	---	---	---	---	---	---	---	---	0.0005	0.067
<i>E_P54NI</i>	---	---	---	---	---	---	---	---	---	0.02
<i>E_P54N2</i>	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are coloured in orange, no significant differences are described by P-values >0.05.

The K_m values of the ‘Satina’ alleles *E_SA* and *E_SNI* differed not significantly but showed a significant difference to *E_SN3*. The allele *E_SN3* revealed the highest K_m of approximately 24mM meaning that this allele shows a lower affinity to its substrate sucrose compared to the other two ‘Satina’ alleles.

K_m values of the two ‘Theresa’ alleles did not differ significantly. The allele *E_TA* had a K_m value of approximately 21mM, whilst the K_m of allele *E_TNI* was around 19mM.

Comparison of K_m values of the two P40 alleles *E_P40NI* and *E_P40N2* revealed no significant change in enzyme affinity to its substrate. K_m values of both enzymes were approximately 17mM. The K_m values of P54 alleles *E_P54NI* and *E_P54N2* differed significantly. *E_P54NI* displayed a K_m value around 23mM and *E_P54N2* of approximately 20mM. The K_m value of the yeast wild type reference strain *FY 1679* was approximately 24mM.

❖ Biochemical characterization of *invGF* alleles

Results of the biochemical analysis for *invGF* alleles are shown in Table 3.2.15

Table 3.2.15: K_m (mM) of *invGF* invertase alleles.

Genotype	<i>invGF</i> allele	K_m
‘Satina’	<i>F_SNI</i>	12.72±1.41
	<i>F_SN2</i>	12.30±1.41
	<i>F_SN3</i>	14.73±1.41
	<i>F_SN4</i>	17.71±1.23
‘Diana’	<i>F_DNI</i>	13.10±1.41
	<i>F_DN2</i>	14.45±1.45
P18	<i>F_P18N</i>	13.90±1.41
P40	<i>F_P40NI</i>	21.41±1.41
	<i>F_P40N2</i>	17.03±1.41
P54	<i>F_P54NI</i>	18.02±1.41
	<i>F_P54N2</i>	16.65±1.41
<i>FY 1679</i>		20.1±3.1

Standard deviations are derived from two biological replicates done in technical triplicates. To make assays of different invertase isoforms comparable, the yeast reference strain *FY 1679* was used as positive control.

Table 3.2.16 summarizes the significance values for differences between the K_m values measured for the *invGF* alleles. The statistical calculation was performed by Benjamin Stich, MPIZ/Köln.

Table 3.2.16: Statistical significance levels of K_m values from the *invGF* invertase alleles.

Allele	<i>F</i> <i>SN1</i>	<i>F</i> <i>SN2</i>	<i>F</i> <i>SN3</i>	<i>F</i> <i>SN4</i>	<i>F</i> <i>DN1</i>	<i>F</i> <i>DN2</i>	<i>F</i> <i>P18N</i>	<i>F</i> <i>P40N1</i>	<i>F</i> <i>P40N2</i>	<i>F</i> <i>P54N1</i>	<i>F</i> <i>P54N2</i>
<i>F</i> _SN1	---	0.83	0.33	0.02	0.87	0.41	0.56	0.0009	0.05	0.02	0.07
<i>F</i> _SN2	---	---	0.24	0.01	0.71	0.31	0.44	0.0006	0.035	0.014	0.05
<i>F</i> _SN3	---	---	---	0.14	0.03	0.1	0.06	0.07	0.72	0.12	0.35
<i>F</i> _SN4	---	---	---	---	0.03	0.11	0.06	0.07	0.72	0.87	0.58
<i>F</i> _DN1	---	---	---	---	---	0.51	0.68	0.001	0.07	0.03	0.1
<i>F</i> _DN2	---	---	---	---	---	---	0.8	0.005	0.23	0.1	0.3
<i>F</i> _P18N	---	---	---	---	---	---	---	0.003	0.14	0.06	0.2
<i>F</i> _P40N1	---	---	---	---	---	---	---	---	0.048	0.11	0.034
<i>F</i> _P40N2	---	---	---	---	---	---	---	---	---	0.63	0.85
<i>F</i> _P54N1	---	---	---	---	---	---	---	---	---	---	0.51
<i>F</i> _P54N2	---	---	---	---	---	---	---	---	---	---	---

Significant differences are defined by P-values <0.05 and are coloured in orange, no significant differences are described by P-values >0.05.

The K_m values of the ‘Satina’ alleles *F*_SN1, *F*_SN2, and *F*_SN3 were similar, whilst the K_m value of allele *F*_SN4 differed significantly. The allele *F*_SN4 showed the lowest substrate affinity compared to the other ‘Satina’ alleles characterized by a K_m value of approximately 18mM. The enzymatic characteristics of the two ‘Diana’ alleles *F*_DN1 (13mM) and *F*_DN2 (≈14mM) were similar to each other, no significant differences were detected.

The K_m values of the P40 alleles *F*_P40N1 and *F*_P40N2 differed significantly. The allele *F*_P40N1 displayed a lower affinity (≈21mM) to sucrose than allele *F*_P40N2 (≈17mM). The K_m values of the two P54 alleles *F*_P54N1 and *F*_P54N2 showed similarity to each other, no enzymatic differences were observed. The K_m value of *F*_P54N1 was 18mM and of *F*_P54N2 approximately 17mM. The K_m value of the yeast wild type reference strain *FY 1679* was approximately 20mM.

❖ Comparison of biochemical *invGE* and *invGF* characteristics

Both *invGE* and *invGF* genes encode cell wall-bound invertase isoforms functional in the apoplast. Biochemical analysis of *invGE* and *invGF* alleles showed differences in enzyme affinity to the substrate sucrose. The significance values for differences between the K_m values measured for *invGE* alleles compared to *invGF* alleles are listed in Table 3.2.17.

Table 3.2.17: Statistical significance levels of K_m values from the *invGE* and *invGF* invertase alleles.

Allele	<i>E</i> <i>SA</i>	<i>E</i> <i>SN1</i>	<i>E</i> <i>SN3</i>	<i>E</i> <i>TA</i>	<i>E</i> <i>TN1</i>	<i>E</i> <i>P18N1</i>	<i>E</i> <i>P40N1</i>	<i>E</i> <i>P40N2</i>	<i>E</i> <i>P54N1</i>	<i>E</i> <i>P54N2</i>
<i>F_SN1</i>	0.002	0.029	4.30 E-05	0.0005	0.005	3.08 E-05	0.026	0.026	0.0003	0.004
<i>F_SN2</i>	0.001	0.019	3.03 E-05	0.0003	0.003	2.19 E-05	0.016	0.017	0.0002	0.003
<i>F_SN3</i>	0.016	0.215	0.0002	0.003	0.043	0.0002	0.192	0.196	0.003	0.049
<i>F_SN4</i>	0.28	0.685	0.0042	0.0669	0.595	0.0028	0.739	0.728	0.1027	0.8082
<i>F_DN1</i>	0.001	0.042	5.71 E-05	0.001	0.0072	4.07 E-05	0.037	0.038	0.001	0.007
<i>F_DN2</i>	0.017	0.205	0.0003	0.004	0.0438	0.0002	0.183	0.187	0.004	0.051
<i>F_P18N</i>	0.007	0.099	0.0001	0.002	0.0179	8.31 E-05	0.087	0.089	0.001	0.018
<i>F_P40N1</i>	0.358	0.025	0.18	0.953	0.1351	0.125	0.029	0.028	0.512	0.051
<i>F_P40N2</i>	0.16	0.985	0.002	0.036	0.3699	0.0015	0.957	0.968	0.049	0.506

Significant differences are defined by P-values <0.05 and are coloured in orange, no significant differences are described by P-values >0.05.

Significant differences in K_m values were detected for the *invGE* and *invGF* alleles of the cultivar ‘Satina’. The alleles *E_SA* and *E_SN1* were significantly different to *F_SN1*, *F_SN2*, and *F_SN3*, whilst *F_SN4* showed no significant differences. The allele *E_SN3* differed significantly from all four *invGF* alleles.

Biochemical analysis of the P18 *invGE* and *invGF* alleles showed significant differences for both alleles. The allele *E_P40N1* converted sucrose with a lower affinity than the allele *F_P40N2*. The *invGF* alleles of P54 possessed a slightly lower affinity to sucrose than the P54 *invGE* alleles.

3.2.2.4 Western blot analysis of *invGE* and *invGF* invertase proteins

To analyze whether the observed differences of invertase activity of *invGE* and *invGF* alleles are due to changes in protein quantity, it is necessary to perform immunoblot quantification. From all invertase antibodies tested (chapter 2, Table 2.2.15), no satisfying *invGE* and *invGF* protein detection pattern were obtained. Therefore, a specific antibody against potato invertases was custom produced by BioGenes, Gesellschaft für Biopolymere mbH (Berlin) and is subject of ongoing investigations of this project.

3.3 The *Invap-a* locus on chromosome X

The *Inv_{ap}-a* locus consists of the two invertase genes *pCD111* and *pCD141*, and encodes cell wall-bound invertase isoforms. These acidic insoluble invertases act in a pH optimum range of 3.5 to 5.1, and are functional in the apoplast (HEDLEY ET AL., 1993, 1994). cDNA of *pCD111* (incomplete sequence; accession: Z21486) and *pCD141* (accession: Z22645) have been cloned and structurally characterized (HEDLEY ET AL. 1993, 1994).

3.3.1 Structural characterization of the genes *pCD111* and *pCD141*

3.3.1.1 Molecular cloning of *pCD111* and *pCD141* invertase cDNA alleles from leaf tissue

Using full-length gene specific primers, cDNA invertase alleles of *pCD111* and *pCD141* were cloned and sequenced from the three tetraploid potato cultivars ‘Satina’, ‘Diana’ and ‘Theresa’, and from the three diploid potato genotypes P18, P40, and P54. The tetraploid genotypes were selected based on the presence of the associated SSCP fragment from the *pCD141* gene (LI ET AL., 2008; Table 3.3.1) and, therefore, were chosen in this study. The diploid potato genotypes were included because they were used as parents for mapping QTLs and candidate genes for cold sweetening of potato tubers (MENÉNDEZ ET AL., 2002).

Table 3.3.1: Distribution of the associated SSCP fragment *pCD141_3c* present in the tetraploid genotypes ‘Satina’, ‘Diana’, and ‘Theresa’.

Genotype	<i>pCD141-3c</i>
‘Satina’	1
‘Diana’	0
‘Theresa’	0

0=SSCP fragment is absent, 1=SSCP fragment is present. Primers used for the amplification of the fragment *pCD141-3* are listed in chapter 2 (Table 2.1.8).

Out of 40 cloned cDNA sequences per genotype, a sequence was defined as an allele when it was detected twice in two independent PCRs⁹. Additionally, the consensus sequence of all alleles found in one genotype was used for allele definition when variable sequence polymorphisms occurred. For the genes *pCD111* and *pCD141* not every allele was detected twice in two independent PCRs due to the focus on the genes *Pain-I* (section 3.1), *invGE*, and *invGF* (section 3.2) because at the beginning of this work less information about association and putative allelic effects on potato chips quality of *pCD111* and *pCD141* was available. *pCD111* and *pCD141* invertase alleles obtained from each genotype are listed in Table 3.3.2.

⁹ Exceptions are listed in Appendix A 3.3.

Table 3.3.2: Overview of *pCD111* and *pCD141* alleles.

Genotype	Full-length clones	<i>pCD111</i> alleles	Full-length clones	<i>pCD141</i> alleles
‘Satina’	5	<i>pCD111_S1</i> <i>pCD111_S2</i> <i>pCD111_S3</i>	6	<i>pCD141_S1</i> <i>pCD141_S2</i> <i>pCD141_S3</i>
‘Diana’	1	<i>pCD111_D1</i>	5	<i>pCD141_D1</i> <i>pCD141_D2</i>
‘Theresa’	4	<i>pCD111_T1</i> <i>pCD111_T2</i>	6	<i>pCD141_T1</i> <i>pCD141_T2</i> <i>pCD141_T3</i>
P18	0	0	5	<i>pCD141_P18_1</i> <i>pCD141_P18_2</i>
P40	1	<i>pCD111_P40_1</i>	2	<i>pCD141_P40_1</i>
P54	3	<i>pCD111_P54_1</i> <i>pCD111_P54_2</i>	4	<i>pCD141_P54_1</i> <i>pCD141_P54_2</i>

The full-length clone number refers to the number of fully sequenced clones from each genotype used for allele definition. PCR amplification was carried out using gene specific full-length primers: *pCD111*-CD111fl_F/CD111fl_R (chapter 2, Table 2.1.2), *pCD141*-CD141fl_F/CD141fl_R (chapter 2, Table 2.1.2).

In ‘Theresa’, P18, and P40 cDNA sequences were found, which contained internal frame shifts and missing or modified start or stop codons in gene *pCD111*. These sequences were excluded from the analysis. From all six genotypes selected in this study, nine *pCD111* and 13 *pCD141* alleles were identified

3.3.1.1.1 *pCD111* cDNA alleles of the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’¹⁰

From five full-length cDNA clones of the genotype ‘Satina’ three different alleles *pCD111_S1*, *pCD111_S2*, and *pCD111_S3* were identified. For the cultivar ‘Diana’ one full-length clone was obtained and named *pCD111_D1*. Cloning and sequencing of ‘Theresa’ cDNA resulted in four full-length clones, from which two *pCD111* alleles *pCD111_T1* and *pCD111_T2* were defined.

Comparing all six alleles at nucleotide level (Appendix A 3.3.6), 47 single nucleotide polymorphisms (SNPs) were detected, 30 of them resulted in an amino acid exchange (Table 3.3.3; Figure 3.3.1).

¹⁰ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: ‘Satina’ (Appendix A 3.3.1), ‘Theresa’ (Appendix A 3.3.2). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *pCD111_S1* (Appendix A 3.3.3), *pCD111_S2* (Appendix A 3.3.4), *pCD111_T2* (Appendix A 3.3.5). For the alleles *pCD111_S3*, *pCD111_D1*, and *pCD111_T1* only one nucleotide sequence was obtained, respectively.

Table 3.3.3: SNPs present in ‘Satina’, ‘Diana’, and ‘Theresa’ *pCD111* alleles.

Position of cDNA SNP	<i>pCD111</i> S1	<i>pCD111</i> S2	<i>pCD111</i> S3	<i>pCD111</i> D1	<i>pCD111</i> T1	<i>pCD111</i> T2	aa
181	A	A	A	A	C	A	C/A Q61N
183	A	T	A	A	A	T	A/T K61N
207	C	C	T	T	C	C	s.
267	T	T	A	A	A	T	s.
303	C	C	G	G	C	C	s.
357	T	C	T	T	T	C	s.
478	A	G	G	G	G	G	A/G N160D
635	G	A	G	G	G	A	G/A G212E
640	C	A	C	C	C	A	C/A Q214K
668	A	G	A	A	A	G	A/G K223R
670	A	G	A	A	A	G	A/G N224D
675	C	C	C	C	A	C	C/A L225F
691	A	G	A	A	A	G	A/G I231V
693	T	C	T	T	T	C	T/C I231V
697	C	G	C	C	C	G	C/G H233D
750	T	C	T	T	T	C	s.
780	G	A	G	G	G	A	s.
815	C	C	T	T	T	C	T/C V272A
858	A	C	A	A	C	C	s.
865	A	A	A	A	C	A	C/A Q289K
866	T	A	T	T	A	A	T/A I289K
897	A	C	A	A	A	C	s.
920	G	T	G	G	G	G	G/T W307
924	G	C	G	G	G	C	G/C K308N
939	C	C	T	T	C	C	s.
972	T	T	C	C	T	T	s.
977	C	T	C	C	C	T	C/T P326L
981	C	G	C	C	C	G	C/G S327R
1039	G	G	A	A	G	G	G/A V347I
1056	A	G	G	G	A	G	s.
1081	T	C	T	T	T	C	s.
1104	G	G	A	A	G	G	s.
1110	G	G	A	A	G	G	s.
1141	C	C	T	T	C	C	C/T L381F
1143	A	A	T	T	A	A	A/T L381F

Position of cDNA SNP	<i>pCD111</i> <i>S1</i>	<i>pCD111</i> <i>S2</i>	<i>pCD111</i> <i>S3</i>	<i>pCD111</i> <i>D1</i>	<i>pCD111</i> <i>T1</i>	<i>pCD111</i> <i>T2</i>	aa
1182	G	C	C	C	C	C	G/C K394N
1245	T	C	C	C	T	C	s.
1252	T	T	T	T	A	T	A/T T418S
1273	C	C	C	C	A	C	A/C T425P
1286	A	A	G	A	A	A	A/G N429S
1292	A	C	C	C	A	C	A/C D431A
1334	C	C	C	C	T	C	T/C M445T
1401	G	A	G	G	G	A	s.
1426	G	G	A	A	G	G	G/A A476T
1432	G	A	G	G	G	A	G/A D478N
1464	A	A	C	C	A	A	s.
1538	T	C	C	C	C	C	T/C I513T

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are orange coloured.

The six *pCD111* alleles from the three tetraploid potato cultivars displayed 47 SNPs resulting in 26 variable amino acid positions because at protein sequence positions 61 and 289 three different amino acids are displayed (Figure 3.3.1). The three alleles of 'Satina' had a total of 41 SNPs, of which 24 caused an amino acid exchange. In the two allelic sequences of 'Theresa' 29 SNPs occurred, from which 19 resulted in amino acid substitutions.

The amino acid alignment shows the polymorphisms of all six *pCD111* alleles from the three tetraploid potato cultivars. The comparison of these deduced protein sequences revealed variable amino acid positions in the different genotypes seen in red, yellow, and grey (Figure 3.3.1).

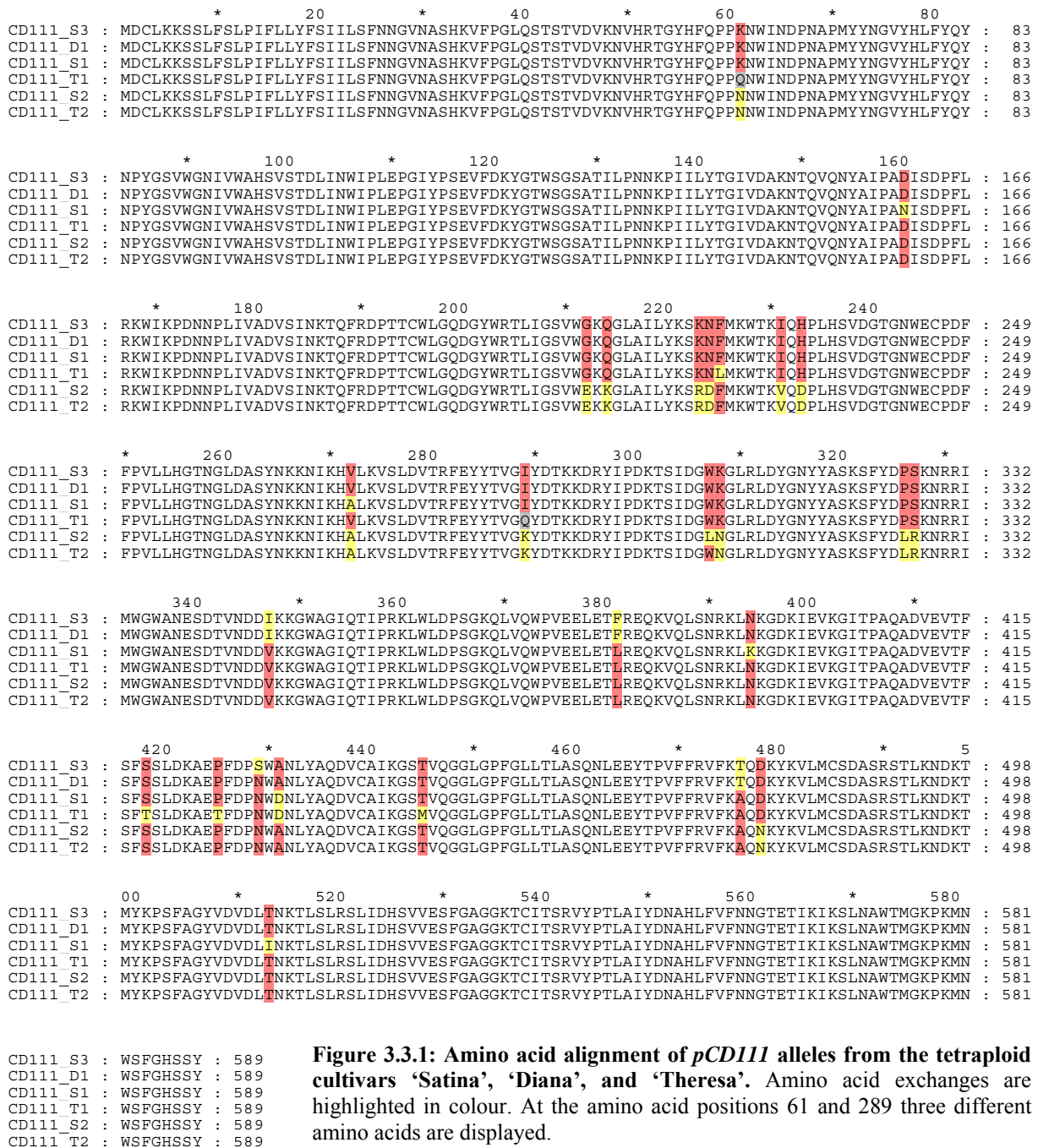


Figure 3.3.1: Amino acid alignment of *pCD111* alleles from the tetraploid cultivars 'Satina', 'Diana', and 'Theresa'. Amino acid exchanges are highlighted in colour. At the amino acid positions 61 and 289 three different amino acids are displayed.

Comparison of allelic cDNA sequences of the gene *pCD111* from the three tetraploid cultivars revealed that none of the six alleles is present in different genotypes.

3.3.1.1.2 *pCD111* cDNA alleles of the diploid potato genotypes P18, P40, and P54¹¹

From the genotype P18 no full-length cDNA alleles were obtained. The cloned sequences either did not feature the start codon of the gene, or the reading frame was shifted resulted in a sequence, which could not be used for further analysis. Cloning and sequencing of P40 cDNA

¹¹ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: P54 (Appendix A 3.3.7). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *pCD111_P54_1* (Appendix A 3.3.8). For the alleles *pCD111_P40_1* and *pCD111_P54_2* only one full-length nucleotide sequence was obtained, respectively.

produced one full-length clone. The allele was named *pCD111_P40_1*. In P54 three full-length clones were found, and two different alleles, *pCD111_P54_1* and *pCD111_P54_2*, were identified.

The nucleotide sequence comparison (Appendix A 3.3.9) of the alleles described above detected 42 SNPs. 22 of them led to amino acid exchanges (Table 3.3.4; Figure 3.3.2).

Table 3.3.4: SNPs present in P40 and P54 alleles.

Position of cDNA SNP	<i>pCD111_P40_1</i>	<i>pCD111_P54_1</i>	<i>pCD111_P54_2</i>	aa
183	A	T	T	A/T K61N
207	T	C	C	s.
267	A	T	T	s.
303	G	C	G	s.
357	T	C	C	s.
360	C	C	T	s.
389	C	C	T	C/T P133L
407	A	A	C	A/C K136T
421	T	T	C	T/C Y141H
633	G	G	C	G/C W211C
635	G	A	A	G/A G212E
640	C	A	A	C/A Q214K
668	A	G	G	A/G K223R
670	A	G	G	A/G N224D
691	A	G	G	s.
693	T	C	C	T/C I231V
697	C	G	G	C/G H233D
750	T	C	C	s.
780	G	A	G	s.
815	T	C	T	C/T A272V
858	A	C	C	s.
866	T	A	A	T/A I289K
897	A	C	C	s.
924	G	C	G	C/G N308K
939	T	C	C	s.
972	C	T	T	s.
977	C	T	T	C/T P326L
981	C	G	G	C/G S327R
1039	A	G	G	A/G I347V
1081	T	C	C	s.
1104	A	G	G	s.
1110	A	G	G	s.

Position of cDNA SNP	<i>pCD111_P40_1</i>	<i>pCD111_P54_1</i>	<i>pCD111_P54_2</i>	aa
1141	T	C	C	s.
1143	T	A	A	T/A F381L
1302	T	T	C	s.
1382	A	A	G	A/G Q461R
1401	G	A	A	s.
1404	G	G	A	s.
1426	A	G	G	A/G T476A
1432	G	A	G	A/G N478D
1464	C	A	A	s.
1757	G	A	A	G/A R586H

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are orange coloured.

In the two P54 allelic sequences *pCD111_P54_1* and *pCD111_P54_2* 10 SNPs occurred. Eight of them caused an amino acid substitution.

The amino acid alignment of the P40 allele *pCD111_P40_1*, and the P54 alleles *pCD111_P54_1* and *pCD111_P54_2* showed 22 polymorphisms. (Figure 3.3.2).

```

CD111_P541 : MDCLKKSSLFSLPIFLLYFSIILSFNNGVNVASHKVFPGQLQSTSTVDVKNVHRTGYHFQPPNNWINDPNAPMYNGVYHLFY : 81
CD111_P542 : MDCLKKSSLFSLPIFLLYFSIILSFNNGVNVASHKVFPGQLQSTSTVDVKNVHRTGYHFQPPNNWINDPNAPMYNGVYHLFY : 81
CD111_P401 : MDCLKKSSLFSLPIFLLYFSIILSFNNGVNVASHKVFPGQLQSTSTVDVKNVHRTGYHFQPPNNWINDPNAPMYNGVYHLFY : 81

CD111_P541 : QYNPYGVSVMGNIVWAHSVSTDINWIPLEPGIYPSEVFDKYGTWSGSATILNNKPIILYTGIVDAKNTQVQNYAIPADIS : 162
CD111_P542 : QYNPYGVSVMGNIVWAHSVSTDINWIPLEPGIYPSEVFDKYGTWSGSATILNNKPIILYTGIVDAKNTQVQNYAIPADIS : 162
CD111_P401 : QYNPYGVSVMGNIVWAHSVSTDINWIPLEPGIYPSEVFDKYGTWSGSATILNNKPIILYTGIVDAKNTQVQNYAIPADIS : 162

CD111_P541 : DPFLRWIKPDNNPLIVADVSINKTQFRDPTTCWLQDGYWRTLGISVWEKKGLAILYKSRDFMKWTKVQDPLHSVDGTGN : 243
CD111_P542 : DPFLRWIKPDNNPLIVADVSINKTQFRDPTTCWLQDGYWRTLGISVCEKKGLAILYKSRDFMKWTKVQDPLHSVDGTGN : 243
CD111_P401 : DPFLRWIKPDNNPLIVADVSINKTQFRDPTTCWLQDGYWRTLGISVWCKQGLAILYKSNFMKWTKYQHPLHSVDGTGN : 243

CD111_P541 : WECPDFFPVLLHGTNGLDASYNKKNIKHVLRKSLDVTREYYTVGKYDTKKDRYIPDKTSIDGWNGLRLDYGNYASKSFY : 324
CD111_P542 : WECPDFFPVLLHGTNGLDASYNKKNIKHVLRKSLDVTREYYTVGKYDTKKDRYIPDKTSIDGWNGLRLDYGNYASKSFY : 324
CD111_P401 : WECPDFFPVLLHGTNGLDASYNKKNIKHVLRKSLDVTREYYTVGKYDTKKDRYIPDKTSIDGWNGLRLDYGNYASKSFY : 324

CD111_P541 : DLRKNRRIMWGWANESDVTNDDVKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETIREQKVQLSNRKLKNGDKIEVKGIT : 405
CD111_P542 : DLRKNRRIMWGWANESDVTNDDVKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETIREQKVQLSNRKLKNGDKIEVKGIT : 405
CD111_P401 : DLRKNRRIMWGWANESDVTNDDVKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETIREQKVQLSNRKLKNGDKIEVKGIT : 405

CD111_P541 : PAQADVEVTFSSSLDKAEPFDPNWNANLYAQDVCAIKGSTVQGGGLGPFGLLTLASQNLLEYTPVFFRVFKAQNKYKVLKMS : 486
CD111_P542 : PAQADVEVTFSSSLDKAEPFDPNWNANLYAQDVCAIKGSTVQGGGLGPFGLLTLASQNLLEYTPVFFRVFKAQNKYKVLKMS : 486
CD111_P401 : PAQADVEVTFSSSLDKAEPFDPNWNANLYAQDVCAIKGSTVQGGGLGPFGLLTLASQNLLEYTPVFFRVFKAQNKYKVLKMS : 486

CD111_P541 : DASRSTLKNKDKTMYKPSFAGYVDVLTNKTLSLRSLIDHSVVSFSGAGGKTCITSRVYPTLAIYDNAHLFVFNNGTETIKI : 567
CD111_P542 : DASRSTLKNKDKTMYKPSFAGYVDVLTNKTLSLRSLIDHSVVSFSGAGGKTCITSRVYPTLAIYDNAHLFVFNNGTETIKI : 567
CD111_P401 : DASRSTLKNKDKTMYKPSFAGYVDVLTNKTLSLRSLIDHSVVSFSGAGGKTCITSRVYPTLAIYDNAHLFVFNNGTETIKI : 567

CD111_P541 : KSLNAWTMGKPKMNWSFGHSSY : 589
CD111_P542 : KSLNAWTMGKPKMNWSFGHSSY : 589
CD111_P401 : KSLNAWTMGKPKMNWSFGHSSY : 589

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Figure 3.3.2: Amino acid alignment of *pCD111* alleles from the diploid genotypes P40 and P54. Amino acid exchanges are highlighted in colour.

3.3.1.1.3 Amino acid alignment of all *pCD111* invertase alleles of the analyzed potato genotypes

Multiple alignment of the nine *pCD111* deduced protein sequences revealed partially overlapping amino acid polymorphisms between the tetraploid and diploid genotypes (Figure 3.3.3). Several amino acid exchanges occur in different genotypes at the same position. The comparison of the protein sequences showed that amino acids differed at 32 positions, of which 17 were genotype specific and occurred only once.

			*		20		*		40		*		60		*		80	
CD111_S3	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_P401	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_D1	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_S1	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_T1	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_S2	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_T2	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_P541	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
CD111_P542	:	MDCLKSSLSFLPIFLLYFSIILSFNNGVNASHKVFPGLQSTSTVDVKNVHRTGYHFQPP																81
			*		100		*		120		*		140		*		160	
CD111_S3	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_P401	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_D1	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_S1	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_T1	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_S2	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_T2	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_P541	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
CD111_P542	:	QYNPYGSVWGNIVWAHSVSTDLINWIPLLEPGIYPSEVFDKYGTWGSATILPNNKPIILYTGIVDAKNTQVQNYAIPADIS																162
			*		180		*		200		*		220		*		240	
CD111_S3	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_P401	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_D1	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_S1	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_T1	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_S2	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_T2	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_P541	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
CD111_P542	:	DPFLRKWKIPDNNPLIVADVSINKTQFRDPTTCWLGDQGYWRTLIGSVWKKQGLAILYKSKNFMKWTKIOHPLHSVDTGTGN																243
			*		260		*		280		*		300		*		320	
CD111_S3	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_P401	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_D1	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_S1	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_T1	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_S2	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_T2	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_P541	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
CD111_P542	:	WECPDFFPVLLHGTNGLDASYNKKNIKHVLKVS LDVTRFEYTVGTYD TKKDRYIPDKTSIDGWLRLDYGNYASKSFY																324
			*		340		*		360		*		380		*		400	
CD111_S3	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_P401	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_D1	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_S1	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_T1	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_S2	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_T2	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_P541	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
CD111_P542	:	DPSKNRRIMWGWANESDTVNDDIKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETFREQKVQLSNRKLKNGDKIEVKGIT																405
			*		420		*		440		*		460		*		480	
CD111_S3	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_P401	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_D1	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_S1	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_T1	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_S2	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_T2	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_P541	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486
CD111_P542	:	PAQADVEVTFSSSLDKAEFPDPNWNALYAQDVCAIKGSTVQGGGLGPFGLLTLASQNL E EYTPVFFRVFKQDKYKVL M C S																486

		*	500	*	520	*	540	*	560					
CD111_S3	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_P401	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_D1	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_S1	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_T1	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_S2	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_T2	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_P541	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567
CD111_P542	:	DASRSTL	KNDK	TMYPKPSFAGYVDV	DL	TNK	TL	SLRSLIDHSV	VFESFGAGGKTCIT	SRVYPTLAIYD	NAHLFVF	NNGTETIKI	:	567

		*	580	
CD111_S3	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_P401	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_D1	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_S1	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_T1	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_S2	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_T2	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_P541	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589
CD111_P542	:	KSLNAWTMGKPKMNWSFG	HSSY	: 589

Figure 3.3.3: Amino acid alignment of all cloned *pCD111* invertase alleles. Amino acid exchanges are highlighted in colour. At the amino acid positions 61 and 289 three different amino acids are displayed.

Figure 3.3.3: Amino acid alignment of all cloned *pCD111* invertase alleles. Amino acid exchanges are highlighted in colour. At the amino acid positions 61 and 289 three different amino acids are displayed.

Comparison of cDNA sequences revealed that the alleles *pCD111_T2* and *pCD111_P54_1* are identical at amino acid level.

3.3.1.1.4 Phenetic trees of all *pCD111* invertase alleles of the analyzed potato genotypes

In addition to the multiple amino acid alignment (3.3.1.1.3), the phenetic tree analysis was used to group the invertase alleles according to similarity. Using the neighbour-joining method showed that *pCD111* alleles from all analyzed potato genotypes grouped in two clades and five subclades (Figure 3.3.4).

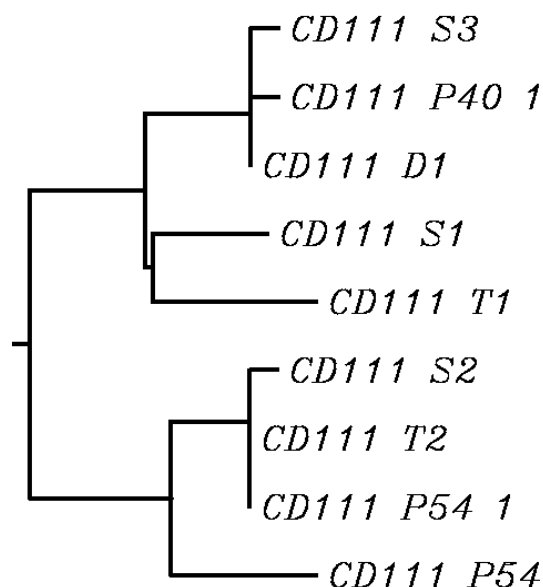


Figure 3.3.4: Amino acid based phenetic tree (Neighbour-joining tree) of all cloned *pCD111* invertase alleles.

The first clade includes alleles *pCD111_S1*, *pCD111_S3*, *pCD111_D1*, *pCD111_T1*, and *pCD111_P40_1*. The second clade consists of *pCD111_S2*, *pCD111_T2*, and the two alleles from the diploid genotype P54 *pCD111_P54_1* and *pCD111_P54_2*. Most alleles of both

clades differ in one or more amino acid positions. The alleles *pCD111_T2* and *pCD111_P54_1* are identical at amino acid level.

Cloning and sequencing of *pCD111* alleles showed that the tetraploid genotype ‘Theresa’ and the diploid genotype P54 contained an allele identical in amino acid sequence but different at nucleotide level. The following Table 3.3.5 summarizes the nucleotide comparison of the two amino acid identical alleles *pCD111_T2* and *pCD111_P54_1*. The allelic nucleotide sequence was defined based on the consensus sequence of the alignment of full-length clones obtained from each genotype (Table 3.3.2). Although SNPs are present at four positions in the cDNAs, the nucleotide polymorphisms resulted in one and the same amino acid sequence.

Table 3.3.5: Genotype specific nucleotide differences of alleles identical at amino acid level.

Position of cDNA SNP	<i>pCD111_T2</i>	<i>pCD111_P54_1</i>
21	C	T
111	A	T
681	A	G
801	G	A

The nucleotide polymorphisms between all *pCD111* (Appendix A 3.3.10) were visualized using the phenetic tree analysis (Figure 3.3.5).

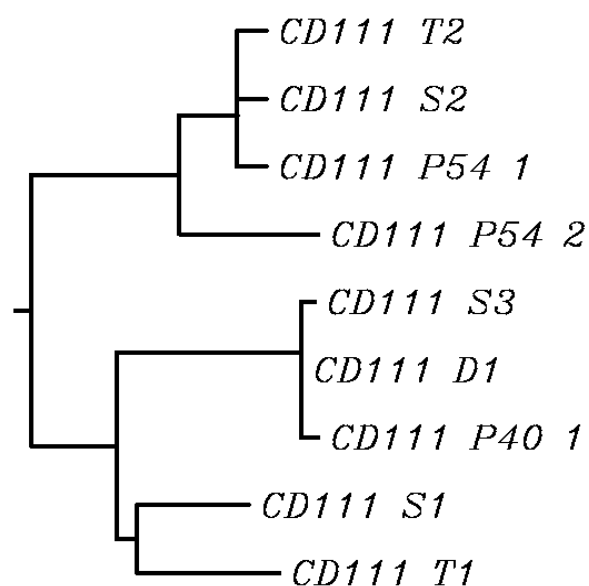


Figure 3.3.5: Nucleotide sequence based phenetic tree (Neighbour-joining tree) of all cloned *pCD111* invertase alleles.

The two phenetic trees are very similar, just displaying more subclades due to a higher number of nucleotide polymorphisms as compared to the amino acid exchanges (Figure 3.3.4).

3.3.1.1.5 *pCD141* cDNA alleles of the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’,¹²

Cloning and sequencing of cDNA of the cultivar ‘Satina’ resulted in six full-length clones, from which three *pCD141* alleles *pCD141_S1*, *pCD141_S2*, and *pCD141_S3* were identified. For the cultivar ‘Diana’ five full-length clones were obtained, and of those two different alleles *pCD141_D1* and *pCD141_D2* were defined. From the cultivar ‘Theresa’ six full-length clones were isolated and led to the definition of three different alleles *pCD141_T1*, *pCD141_T2*, and *pCD141_T3*.

At nucleotide level (Appendix A 3.3.19) the alleles contain 62 SNPs but not all of them are causative for amino acid exchanges (Table 3.3.6; Figure 3.3.6).

Table 3.3.6: SNPs present in ‘Satina’, ‘Diana’, and ‘Theresa’ *pCD141* alleles.

Position cDNA SNP	<i>pCD141</i> <i>S1</i>	<i>pCD141</i> <i>S2</i>	<i>pCD141</i> <i>S3</i>	<i>pCD141</i> <i>D1</i>	<i>pCD141</i> <i>D2</i>	<i>pCD141</i> <i>T1</i>	<i>pCD141</i> <i>T2</i>	<i>pCD141</i> <i>T3</i>	aa
57	C	T	C	C	C	T	C	C	s.
140	C	T	C	C	C	T	C	C	T/C V47A
178	C	C	T	C	C	C	C	C	C/T P60S
186	C	T	C	C	C	T	C	C	s.
207	T	T	T	C	C	T	C	C	s.
276	C	T	T	T	T	T	T	T	s.
280	A	G	G	G	G	G	G	G	A/G I94V
378	C	C	C	T	T	C	T	T	s.
426	C	T	T	T	T	C	T	T	s.
440	C	C	C	G	G	C	G	G	C/G A147G
444	T	C	C	T	T	T	T	T	s.
462	T	C	T	C	C	C	C	C	s.
474	G	A	G	G	G	G	G	G	s.
483	G	A	G	G	G	G	G	G	G/A M161I
508	A	A	A	A	A	A	A	G	A/G I170V
582	A	A	A	C	C	A	C	C	s.
601	G	G	A	G	G	G	G	G	G/A G201R
621	T	T	T	A	A	A	A	A	s.
624	G	G	G	C	C	C	C	C	s.
667	A	G	A	A	A	A	A	A	A/G N223D

¹² Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: ‘Satina’ (Appendix A 3.3.11), ‘Diana’ (Appendix A 3.3.12), ‘Theresa’ (Appendix A 3.3.13). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *pCD141_S2* (Appendix A 3.3.14), *pCD141_S3* (Appendix A 3.3.15), *pCD141_D1* (Appendix A 3.3.16), *pCD141_T1* (Appendix A 3.3.17), *pCD141_T2* (Appendix A 3.3.18). For the alleles *pCD141_S1*, *pCD141_D2*, and *pCD141_T3* only one full-length nucleotide sequence was obtained, respectively.

Position cDNA SNP	<i>pCD141</i> <i>S1</i>	<i>pCD141</i> <i>S2</i>	<i>pCD141</i> <i>S3</i>	<i>pCD141</i> <i>D1</i>	<i>pCD141</i> <i>D2</i>	<i>pCD141</i> <i>T1</i>	<i>pCD141</i> <i>IT2</i>	<i>pCD141</i> <i>T3</i>	aa
673	A	G	A	A	A	A	A	A	A/G N225D
714	A	C	A	A	A	A	A	A	s.
717	T	C	C	C	C	C	C	C	s.
720	C	G	C	C	C	C	C	C	s.
765	A	G	A	A	A	A	A	A	s.
775	G	A	G	G	G	G	G	G	G/A D259N
798	T	T	T	C	C	C	C	C	s.
843	T	T	T	A	A	A	A	A	s.
862	A	G	A	A	A	A	A	A	A/G I288V
889	A	A	A	C	C	C	C	C	s.
891	G	A	A	G	G	G	G	G	s.
892	C	T	T	T	T	T	T	T	C/T H298Y
905	A	A	A	G	G	G	G	G	A/G N302S
939	T	T	T	C	C	C	C	C	s.
980	G	A	G	G	G	G	G	G	G/A S327N
1029	T	T	G	T	T	T	T	T	s.
1030	C	G	G	G	G	G	G	G	C/G R344V
1031	G	T	T	T	T	T	T	T	G/T R344V
1059	C	T	C	C	C	C	C	C	s.
1096	T	C	C	C	C	C	C	C	T/C S366P
1135	T	T	C	T	T	T	T	T	s.
1188	G	A	G	G	G	G	G	G	s.
1224	A	T	A	A	A	A	A	A	s.
1255	G	A	A	A	A	A	A	A	G/A A419T
1266	A	T	T	C	C	C	C	C	A/T E422D
1277	T	C	C	C	C	C	C	C	T/C L426P
1368	C	C	C	A	A	A	A	A	s.
1416	T	T	T	C	C	C	C	C	s.
1434	A	A	A	T	T	T	T	T	A/T Q478H
1446	G	A	A	A	A	A	A	A	s.
1467	C	T	T	C	C	C	C	C	s.
1503	T	C	C	T	T	T	T	T	s.
1541	C	C	C	C	T	C	C	C	C/T A514V
1542	A	G	G	G	G	G	G	G	s.
1560	T	T	T	C	C	C	C	C	s.
1582	G	G	G	A	A	A	A	A	G/A V528I
1614	G	A	A	A	A	A	A	A	s.
1629	G	G	G	A	A	A	A	A	s.
1641	A	G	G	G	G	G	G	G	s.
1674	C	A	A	A	A	A	A	A	s.
1680	C	C	T	C	C	C	C	C	s.
1683	T	C	C	C	C	C	C	C	s.


```

00          *          520          *          540          *          560          *          580
CD141_T2 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_T3 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_D1 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_D2 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_T1 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_S2 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_S3 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581
CD141_S1 : TMYKPSFAGYVDVDLADKKLSLRSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITETLNAWSMANAKL : 581

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CD141_T2 : H : 582
CD141_T3 : H : 582
CD141_D1 : H : 582
CD141_D2 : H : 582
CD141_T1 : H : 582
CD141_S2 : H : 582
CD141_S3 : H : 582
CD141_S1 : H : 582

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Figure 3.3.6: Amino acid alignment of *pCD141* alleles from the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’. Amino acid exchanges are highlighted in colour.

Comparison of allelic amino acid sequences from the tetraploid cultivars revealed that the genotypes ‘Diana’ and ‘Theresa’ harbour an allele identical at amino acid level (*CD141_D1*=*CD141_T2*).

3.3.1.1.6 *pCD141* cDNA alleles of the diploid potato genotypes P18, P40, and P54¹³

From the genotype P18 five full-length clones were obtained, from which two different cDNA alleles *pCD141_P18_1* and *pCD141_P18_2* were identified. Cloning and sequencing of P40 cDNA resulted in two full-length clones identical at amino acid level. The allele was named *pCD141_P40_1*. Sequence comparison suggested a second P40 allele but the clones harbouring an allelic sequence either did not feature the start codon of the gene or the reading frame was shifted, and, therefore, were not used in further analysis. From the genotype P54 four full-length clones were isolated, from which two different alleles *pCD141_P54_1* and *pCD141_P54_2* were defined.

The alleles contain at nucleotide level 64 SNPs (Appendix A 3.3.28). These sequence polymorphisms can cause amino acid exchanges (Table 3.3.7; Figure 3.3.7).

Table 3.3.7: SNPs present in P18, P40, and P54 *pCD141* alleles.

Position of cDNA SNP	<i>pCD141_P18_1</i>	<i>pCD141_P18_2</i>	<i>pCD141_P40_1</i>	<i>pCD141_P54_1</i>	<i>pCD141_P54_2</i>	aa
57	T	C	C	C	C	s.
101	T	T	T	T	C	T/C A34V
140	T	C	C	C	C	T/C V47A
141	T	T	T	T	A	s.
153	T	T	T	C	T	s.

¹³ Alleles of each genotype were defined by alignment of all fully sequenced cDNA clones: P18 (Appendix A 3.3.20), P40 (Appendix A 3.3.21), P54 (Appendix A 3.3.22). The nucleotide sequence for one allele was defined by aligning all obtained nucleotide sequences for this allele and using the consensus sequence when SNPs occurred: *pCD141_P18_1* (Appendix A 3.3.23), *pCD141_P18_2* (Appendix A 3.3.24), *pCD141_P40_1* (Appendix A 3.3.25), *pCD141_P54_1* (Appendix A 3.3.26), *pCD141_P54_2* (Appendix A 3.3.27). For the allele *pCD141_P40_2* only one full-length nucleotide sequence was obtained.

Position of cDNA SNP	<i>pCD141</i> <i>P18_1</i>	<i>pCD141</i> <i>P18_2</i>	<i>pCD141</i> <i>P40_1</i>	<i>pCD141</i> <i>P54_1</i>	<i>pCD141</i> <i>P54_2</i>	aa
186	T	C	C	C	C	s.
195	C	C	C	T	C	s.
207	C	T	C	C	T	s.
231	T	C	T	T	C	s.
276	T	C	T	T	C	s.
378	C	C	T	C	C	s.
440	C	C	G	C	C	G/C G147A
444	C	C	T	T	T	s.
462	T	T	C	C	C	s.
582	A	C	C	C	C	s.
585	A	T	A	T	T	s.
621	T	A	A	T	T	s.
624	G	G	C	G	G	s.
701	A	T	A	A	A	A/T H234I
714	G	G	A	A	A	s.
717	C	C	C	C	T	s.
720	G	C	C	G	G	s.
761	C	C	C	C	T	C/T S254L
765	G	A	A	G	G	s.
775	A	G	G	A	A	A/G N259D
798	T	T	C	T	T	s.
843	T	A	A	A	A	s.
862	G	A	A	A	A	G/A V288I
889	A	A	C	A	A	C/A K297Q
891	A	G	G	A	A	s.
895	T	A	T	T	T	T/A F299I
905	A	A	G	A	A	A/G N302S
913	A	A	A	A	G	A/G I305V
939	T	T	C	T	T	s.
980	A	G	G	G	G	A/G N327S
1043	T	T	T	T	C	T/C V348A
1143	A	T	T	T	T	s.
1146	G	A	A	A	A	s.
1158	A	G	G	G	G	s.
1188	A	A	G	G	G	s.
1192	G	G	G	A	A	G/A E398K
1224	A	T	A	A	A	s.
1266	T	T	C	T	T	s.
1368	A	C	A	C	C	s.
1390	T	G	T	T	T	T/G L464V
1412	T	T	T	T	C	T/C F417S
1416	C	T	C	T	C	s.

Position of cDNA SNP	<i>pCD141 P18_1</i>	<i>pCD141 P18_2</i>	<i>pCD141 P40_1</i>	<i>pCD141 P54_1</i>	<i>pCD141 P54_2</i>	aa
1417	A	A	A	A	C	s.
1429	A	G	G	G	G	A/G T477A
1434	T	A	T	T	T	T/A H478Q
1446	G	A	A	G	G	s.
1452	T	T	T	T	C	s.
1461	C	C	C	T	C	s.
1467	C	T	C	C	C	s.
1503	T	C	T	T	T	s.
1542	A	G	G	G	G	s.
1560	T	T	C	T	T	s.
1582	G	G	A	G	G	G/A V528I
1587	G	G	G	C	G	s.
1602	T	T	C	C	T	s.
1613	C	C	C	C	T	C/T T538I
1629	G	G	A	G	G	s.
1641	G	G	G	A	G	s.
1673	C	C	C	T	C	T/C V558A
1674	A	A	A	G	A	s.
1683	C	C	C	C	T	s.
1689	G	G	G	A	G	s.

SNP position numbering refers to cDNA sequence where '1' represents the adenine of the start codon ATG and the polymorphism is described. aa=amino acid exchanges ; s.=synonymous nucleotide exchange that does not lead to an amino acid exchange. Non synonymous nucleotide exchanges lead to an amino acid exchange and are orange coloured.

The five alleles obtained from the three diploid genotypes differed in 64 SNPs, from which 21 resulted in an amino acid exchange. The two P18 alleles differed in 30 SNPs. Nine of them led to an amino acid substitution. In the two P54 allelic sequences 18 SNPs occurred, from which four resulted in an amino acid exchange.

Multiple alignment of the protein sequences of all alleles from the three potato genotypes P18, P40, and P54 showed the variability of positions where amino acids differed (Figure 3.3.7).

Comparison of allelic cDNA sequences of the gene *pCD141* from the three diploid genotypes revealed that none of the five alleles is present in different genotypes.

CD141_S1	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKHFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_S3	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_P182	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYIPDNTSIDGWKGLRLDYGNYA	:	320
CD141_T2	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_T3	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_D1	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_D2	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_P401	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_T1	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_S2	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_P181	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_P541	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320
CD141_P542	:	GTGNWECPDFFPVSLKNDGLDTSYNGDKIKHVLKVSFDVTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYA	:	320

CD141_S1	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_S3	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_P182	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_T2	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_T3	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_D1	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_D2	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_P401	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_T1	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_S2	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_P181	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_P541	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400
CD141_P542	:	SKTFFDSGKNRRILLGWANESDTRDNDVRKGWAGVHPIPRKIWLDSGKQLVQWPVQELETLRKKKVQLNNKKLNKGEKV	:	400

CD141_S1	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_S3	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_P182	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_T2	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_T3	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_D1	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_D2	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_P401	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_T1	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_S2	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_P181	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_P541	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480
CD141_P542	:	EIKGITVAQADVEVIFSFLSLEKAEFFDPSWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDK	:	480

CD141_S1	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_S3	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_P182	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_T2	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_T3	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_D1	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_D2	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_P401	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_T1	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_S2	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_P181	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_P541	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560
CD141_P542	:	YKVLMCSDASRSSLKNETTMYKPSFAGYVDVLDADKKLSLRSLIDHSSVESFGAGGKTCTITSRVYPTLAIFDKAHLFAFN	:	560

CD141_S1	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_S3	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_P182	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_T2	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_T3	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_D1	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_D2	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_P401	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_T1	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_S2	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_P181	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_P541	:	NGAERITITETLNAWSMANAKLH	:	582
CD141_P542	:	NGAERITITETLNAWSMANAKLH	:	582

Figure 3.3.8: Amino acid alignment of all cloned *pCD141* invertase alleles. Amino acid exchanges are highlighted in colour.

sequences revealed that *pCD141_P40_1* differs in one SNP at cDNA position 1602. The P40 allele contains at the given position nucleotide C, whilst *pCD141_D1* and *pCD141_T2* exhibit the nucleotide T.

The nucleotide polymorphisms between all *pCD141* (Appendix A 3.3.29) were visualized using the phenetic tree analysis (Figure 3.3.10).

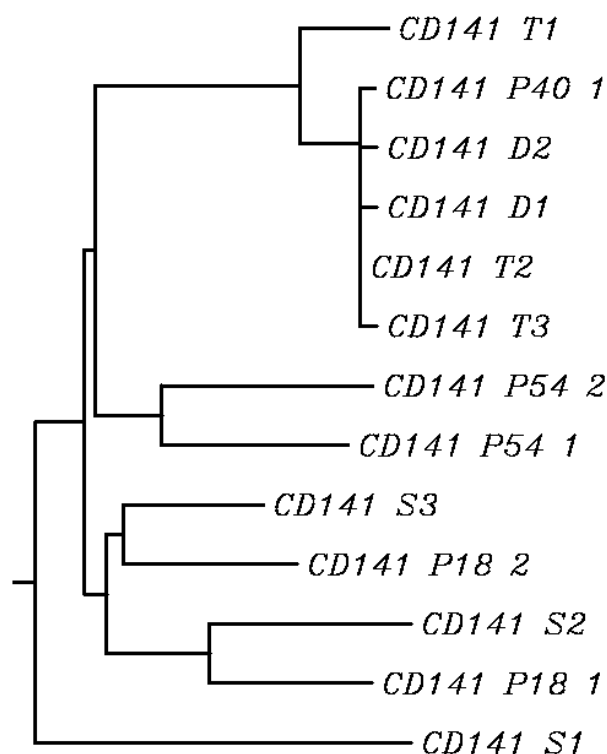


Figure 3.3.10: Nucleotide sequence based phenetic tree (Neighbour-joining tree) of all cloned *pCD141* invertase alleles.

The alleles group regarding their similarity in the same clades and subclades as observed in the amino acid sequences based phenetic tree (Figure 3.3.9). Subclades are more subdivided because of nucleotide polymorphisms.

3.3.1.2 Classification of alleles of the genes *pCD111* and *pCD141*

The genes *pCD111* and *pCD141* map to potato chromosome X (CHEN ET AL., 2001) in a region associated with tuber quality traits where a QTL for potato tuber sugar content, *Sug10a*, was identified (MENÉNDEZ ET AL., 2002). In the latter study, the genes *pCD111* and *pCD141* (referred to *Inv_{ap-a}*) were directly mapped, and showed linkage to the QTL *Sug10a*.

Single-strand conformation polymorphism (SSCP) analysis was carried out for the gene *pCD141* and revealed an association of *pCD141* SSCP fragments with starch and sugar content of potato tubers (LI ET AL., 2008). The SSCP fragment *pCD141_3c* was found to have a negative effect on potato chips quality and tuber starch content. To assign one of the cloned

pCD141 allele (section 3.3.1.1.5/6) to the corresponding associated SSCP fragment, 12 potato cultivars from BNA, SARA, and NOR, respectively were analyzed. To date in none of these 36 genotypes, defined as standards and characterized regarding *pCD141_3c* distribution, associated SNPs could be identified. This is a subject of ongoing investigations in the research project.

3.3.1.3 Genomic organization of the *Invap-a* locus

❖ BAC library screens

The genomic sequence and gene organization of the gene pair *pCD111/pCD141* were identified using high density BAC library screens with two different PCR generated probes. Using the primers CD111S2_F/CD111S1_Rev and pCD141-3F/pCD141-3R (chapter 2, Table 2.1.10A), Probe 1 for the gene *pCD111* and Probe 2 for the gene *pCD141* were generated. Both probes consisted of exon based sequences (HEDLEY ET AL. 1993, 1994) of the genes *pCD111* and *pCD141*, respectively.

The screening of two different BAC libraries (BA and BC, BALLVORA ET AL. 2002, 2007) constructed with genomic DNA of the same diploid genotype resulted in six positive BAC clones for the gene *pCD141* (Table 3.3.8). From the *pCD141* positive BACs one clone (BAC BC3) showed full-length PCR amplification of the gene *pCD111*. BAC screens with the specific *pCD111* Probe 1 were negative.

Table 3.3.8: Positive BAC clones for the genes *pCD111* and *pCD141*.

Library	BAC clones <i>pCD141</i>	PCR <i>pCD141</i>	PCR <i>pCD111</i>
BA BAC library	BA1: Plate 28K15	yes	not analyzed
	BA2: Plate 40M19	yes	not analyzed
	BA3: Plate 59E10	yes	not analyzed
BC BAC library	BC1: Plate 37C23	yes	no
	BC2: Plate 146K10	yes	no
	BC3: Plate 163L15	yes	no

PCR for the genes *pCD111* and *pCD141* were performed with gene specific full-length gene primers: *pCD111*-CD111fl_F/CD111fl_R (chapter 2, Table 2.1.2), *pCD141*-CD141fl_F/CD141fl_R (chapter 2, Table 2.1.2). Positive BACs are numbered and their position in the *E. coli* microtiter plate is listed.

BAC library screening and full-length PCR amplification of *pCD111* and *pCD141* within BAC BC3 suggested that this clone contains both invertase genes of interest. BAC insert (size: 130kb) sequencing was carried out by MWG Biotech AG, Ebersberg using the 454 sequencing technique on the GS FLX system.

❖ BAC Annotation

Full-length sequencing of the BAC insert revealed sequence and structural information of the genes *pCD111* and *pCD141* and of flanking genes (Table 3.3.9).

Table 3.3.9: BC3 sequence annotation.

Strand	Apollo name	Position on the BAC insert (bp)	Description
+	gene 10	2,068-2454	response to auxin stimulus
+	gene 20	4,655-13,035	kinesin
+	gene 30	31,961-36,971	cell wall invertase <i>pCD111</i>, beta-fructofuranosidase (glycoside hydrolase family 32)
+	gene 40	44,311-48,793	cell wall invertase <i>pCD141</i>, beta-fructofuranosidase (glycoside hydrolase family 32)
+	gene 50	62,479-64,535	GTP-binding protein
+	gene 60	69,953-71,012	unknown
+	gene 70	73,520-74,558	putative integral membrane family protein
+	gene 80	75,343-78,806	putative RNA-binding protein
+	gene 90	85,626-88,464	putative embryo defective protein
+	gene 100	122,735-125,703	unknown
-	gene 110	54,022-51,302	putative ribosomal protein
-	gene 120	58,824-55,722	putative esterase lipase
-	gene 130	68,349-66,377	putative dynamin
-	gene 140	81,608-80,442	unknown
-	gene 150	94,506-89,268	DNA-binding protein
-	gene 160	109,512-106,330	Pre-mRNA splicing factor

The invertase genes were named ‘gene 30’ (*pCD111*) and ‘gene 40’ (*pCD141*) in the Apollo BAC sequence characterization. BAC annotation was carried out using the software Apollo Genome Annotation and Curation Tool, version 1.9.8. Both invertase genes are written in bold.

The screened BAC libraries BA and BC harbour genomic DNA of the same diploid genotype P6/210, which is a hybrid derived from the cross of the parental genotypes P40 x P41 (LEISTER ET AL., 1996). The genotype P40 was also selected in this study for invertase allele characterization. Sequence alignments of P40 cDNA alleles from *pCD111* and *pCD141* with the genomic sequences of the genes 30 (*pCD111*, Appendix A 3.3.30) and 40 (*pCD141*, Appendix A 3.3.31) from the BAC BC3 showed no sequence identity. Since only one P40 cDNA allele from *pCD111* and *pCD141* respectively, was cloned it is possible that the second not detected P40 allele corresponds to the BAC sequences, or that the detected BAC allele for the genes *pCD111* and *pCD141* originate from the other parental genotype P41.

❖ Structural characterization of the genes *pCD111* and *pCD141*

The exon and intron organization of the genes *pCD111* and *pCD141* was determined by aligning the *pCD111* (Appendix A 3.3.32¹⁴) and *pCD141* (Appendix A 3.3.33¹⁵) cDNA alleles and the genomic sequences of the corresponding genes from BAC BC3. Both genes consist of six exons and five introns (Figure 3.3.11). The gene *pCD111* has a length of 5012bp, whilst the gene size of *pCD141* is 4478bp.

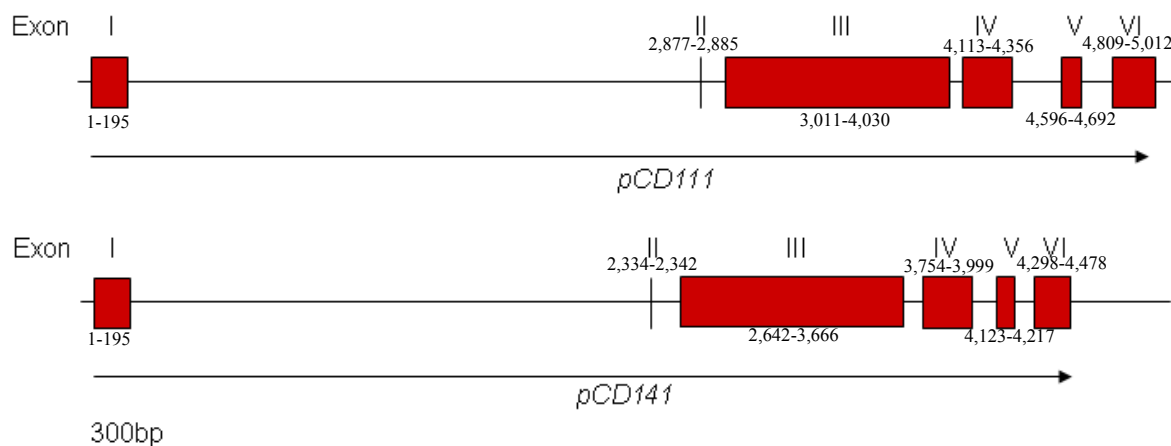


Figure 3.3.11: Genomic organization of the genes *pCD111* and *pCD141*. Exons are drawn in red and numbered from I to VI. The arrows symbolize the whole length of the particular gene without promoter and terminator sequences.

The range of individual *pCD111* and *pCD141* exons and introns are summarized in Table 3.3.10.

Table 3.3.10: Ranges of exons and introns of the genes *pCD111* and *pCD141*.

Exon number	Range (bp)	Intron number	Range (bp)
<i>pCD111</i>			
I	1-195	I	196-2,876
II	2,877-2,885	II	2,886-3,010
III	3,011-4,030	III	4,031-4,112
IV	4,113-4,356	IV	4,357-4,595
V	4,596-4,692	V	4,693-4,808
VI	4,809-5,012		
<i>pCD141</i>			
I	1-195	I	196-2,333
II	2,334-2,342	II	2,343-2,641
III	2,642-3,666	III	3,667-3,753
IV	3,754-3,999	IV	4,000-4,122
V	4,123-4,217	V	4,218-4,297
VI	4,298-4,478		

¹⁴ The alignment program Multalin version 5.4.1 (<http://bioinfo.genetoul.fr>) used revealed problems in the comparison of the genomic *pCD141* sequence with the *pCD141* cDNA sequence due to SNPs in both sequences. The mini-exon II was not aligned properly, two additional nucleotides occurred, whilst another one was connected with the first exon.

¹⁵ As described above in footnote 14.

It is known from the literature that the genomic structure of higher plant invertases is fairly conserved and consists of six to eight exons. As it is the case for the invertases *Pain-1* (chapter 3.1), *invGE* and *invGF* (chapter 3.2), also the genes *pCD111* and *pCD141* show this exon-intron structure and exhibit the extremely small exon II, which only codes for the core tripeptide DPN of the conserved β -fructosidase motif NDPNG (TYMOWSKA-LALANNE & KREIS, 1998).

It has been shown that potato invertase loci encoding cell wall-bound isoforms are characterized by a direct tandem repeat organization where two linked genes are separated by about 2,3kb (MADDISON ET AL., 1999). The *Inv_{ap}-a* locus consisting of the gene pair *pCD111/pCD141* showed this kind of genomic structure with approximately 8kb separating the corresponding invertase genes (Figure 3.3.12).

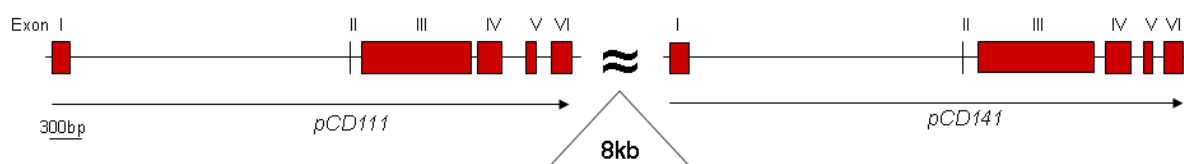


Figure 3.3.12: Tandem repeat linkage of the genes *pCD111* and *pCD141*. Exons are drawn in red and numbered from I to VI. The arrows symbolize the whole length of the genes without promoter and terminator sequences.

4 Discussion¹⁶

4.1 The physiological impact of potato invertases on tuber chips quality

The interest of this study was to characterize the underlying mechanisms of the trait ‘potato chips quality’, which is influenced by the starch and sugar content of potato tubers. A high content of the reducing sugars glucose and fructose, which are accumulating during tuber cold storage (cold-sweetening), results in inferior chips quality e.g. a dark colour, a bitter taste and a high acrylamide concentration. Starch and sugar content of potato tubers are quantitative traits, which can be considered as model traits using the candidate gene approach to unravel the molecular basis of quantitative trait loci (QTL). QTL for tuber starch and sugar content or potato chips colour have been mapped in potato (DOUCHES & FREYRE, 1994; MENÉNDEZ ET AL., 2002). A number of candidate genes have been identified regarding the fact of their co-localisation with QTLs on molecular maps, as well as being functional in the biosynthesis, degradation, or transport of starch and sugars in potato and other plants (CHEN ET AL., 2001; MENÉNDEZ ET AL., 2002). Among others, invertase genes were identified as positional candidates for cold-sweetening QTLs. Furthermore, association analysis showed a significant correlation between alleles of vacuolar and cell wall-bound invertase isoforms and better potato chips quality as well as other tuber traits (LI ET AL., 2005, 2008). In potato three invertase loci *Pain-1*, *Inv_{ap}-a* and *Inv_{ap}-b* are known. The gene *Pain-1* on chromosome III encodes a vacuolar invertase. The two gene pairs *invGE/invGF* of the locus *Inv_{ap}-b* and *pCD111/pCD141* of the locus *Inv_{ap}-a* on potato chromosomes IX and X respectively, code for cell wall-bound invertase isoforms. The genes *invGE* and *invGF* are linked in a direct tandem repeat and separated by approximately 2.3kb from each other. The size of the *Inv_{ap}-b* locus is approximately 8.6kb (MADDISON ET AL., 1999). Comparative analysis of the potato genetic map revealed that the gene pair *pCD111/pCD141* on chromosome X arose by partial chromosome duplication of the gene pair *invGE/invGF* of chromosome IX and, therefore, might also be organized in a direct tandem repeat (GEBHARDT ET AL., 2003; FRIDMAN ET AL., 2003). The present study showed that the genes *pCD111/pCD141* are arranged in a direct tandem repeat and separated by approximately 8kb. The size of the *Inv_{ap}-a* locus is approximately 17.5kb (chapter 3.3, section 3.3.1.3).

¹⁶ Due to the complexity of the present work, a summary of the project and the main results were included to facilitate reading and understanding.

Besides being positional candidates, invertases are also functional candidates. Invertases are enzymes, which catalyze the last step in the carbohydrate breakdown chain. The products of the invertase reaction are the reducing sugars glucose and fructose, which directly interfere with potato chips quality (SHALLENBERGER ET AL., 1959). In consequence invertases fulfil both criteria, being candidate genes in the genetic and biochemical sense.

Part of the phenotypic differences of potato chips quality can be explained by allelic variation of invertases as identified by association analysis (LI ET AL., 2005, 2008). FRIDMAN ET AL. (2004) identified the tomato invertase gene *LIN5* as causal for the QTL *Brix9-2-5* for sugar yield of tomato fruits, comparing differences between *LIN5* alleles of the cultivated tomato (*Solanum lycopersicum*) and of wild species (*Solanum pennellii*). The tomato fruits of the wild species showed a dramatic reduction of the sugar content compared to fruits of the cultivated tomato. The present study aimed to elucidate whether allelic variation of functional potato invertase alleles in different genotypes can be identified and to what extent this natural variation accounts for the observed phenotypic diversity regarding chips quality. The question was whether allelic variation manifested itself at functional level as functional differences that could be characterized, like observed in tomato for the fruit sugar content (FRIDMAN ET AL., 2004).

To analyze allelic composition and functional relevance of invertase alleles, six different genotypes were selected. Based on association analysis the tetraploid cultivars ‘Satina’, ‘Diana’, and ‘Theresa’ were chosen due to the presence or absence of associated invertase SSCP fragments (LI ET AL., 2005, 2008). Additionally, three diploid potato genotypes were analyzed previously being used as parents for mapping QTLs and candidate genes for cold sweetening of potato tubers (MENÉNDEZ ET AL., 2002).

4.2 Structural characterization of potato invertase alleles

Invertase alleles of the five genes *Pain-1*, *invGE/invGF*, and *pCD111/pCD141* were structurally analyzed focusing on their cDNA and genomic sequences, their exon/intron structures and their gene organization. Additionally, *Pain-1* and *invGE/invGF* alleles were characterized using 3D-modelling to visualize putative effects of allelic amino acid differences. The genes *pCD111* and *pCD141* were not modelled because less information about association and putative allelic effects on potato chips quality was available at the beginning of this work.

❖ Molecular cloning of invertase cDNA alleles

cDNA alleles of *Pain-1* were isolated from tuber tissue, *invGE*, *pCD111*, and *pCD141* cDNA alleles were obtained from leaf tissue. In the case of *invGF*, alleles were isolated using floral as well as leaf tissue. In the following Table 4.1 all invertase cDNA alleles cloned from the six genotypes are listed.

Table 4.1: Overview of isolated invertase cDNA alleles.

Genotype	<i>Pain-1</i> alleles	<i>invGE</i> alleles	<i>invGF</i> alleles	<i>pCD111</i> alleles	<i>pCD141</i> alleles
‘Satina’	2	4	4	3	3
‘Diana’	3	3	2	1	2
‘Theresa’	2	4	2	2	3
P18	2	2	1	0	2
P40	2	2	2	1	1
P54	1	2	2	2	2

PCR based cDNA cloning of the five known potato invertase genes *Pain-1* on chromosome III, *invGE/invGF* on chromosome IX and *pCD111/pCD141* on chromosome X resulted in 64 distinct alleles.

Specific PCR amplification of the five invertase genes *Pain-1*, *invGE*, *invGF*, *pCD111*, and *pCD141* was indicated by the presence of corresponding alleles amplified in one course of PCR using a given gene specific primer pair. In Table 4.2 the sequences of full-length primers for all five genes are listed showing gene specific sequence polymorphisms that resulted in gene specific amplification.

Table 4.2: Full-length primers used for gene specific PCR amplification.

Invertase gene	Chr. number	Forward primer (in 5'-3'orientation)	Reverse primer (in 5'-3'orientation)
<i>Pain-1</i>	III	ATG GCC ACG CAG TAC C	GAT GAA TTA CAA GTC TTG CAA GGG
<i>invGE</i>	IX	ATG GAA TTA TTT ATG AAA AGC TCT TCT CTT TGG GGG T	TTA GTG CAT CTT AGG TAC ATC CAT GCT CCA AGC
<i>invGF</i>	IX	ATG GAT TAT TCA TCT AAT TCT CGT TGG GCT TTG CCA G	TCA ATA TTG TAT CTT AGC TTT GCC CAT ACT CCA TGC
<i>pCD111</i>	X	ATG GAT TGT TTA AAA AAG TCT TCT C	TCA ATA AGA AGA GTG ACC AAA TGA CCA ATT CA
<i>pCD141</i>	X	ATG GAG ATT TTA AGA AGA TCT TCT TCT CTT TGG GTT	CTA GTG CAA CTT TGC ATT AGC CAT GCT CCA AGC

In the course of cloning and sequencing of cDNA alleles, quite a number of singular nucleotide variants (singletons) or non functional clones showing internal frame shifts and missing start and stop codons, were isolated. As the cloning of cDNA alleles was PCR based, a proofreading *Taq*-Polymerase, which is characterized by a high fidelity (approximately 5.5×10^{-5} mismatches per base pair per PCR cycle) was used to minimize PCR derived errors. Hereby four times better amplification results compared to common *Taq* DNA-Polymerase were achieved. Sequence reliability was similar to other commercially available proof-reading polymerases (i.e. *Pfu* DNA-Polymerase: 1.5×10^{-5} errors per base pair per PCR cycle, Invitrogen, Karlsruhe). Another source of *in-vitro* invertase sequence variability might have been the conversion of RNA to cDNA by Superscript II (Invitrogen, Karlsruhe), which possesses an error rate of 1×10^{-4} per nucleotide.

Randomly occurring sequencing errors were eliminated by checking forward and reverse sequence of the affected area. Furthermore, in the present study PCR did not show equal amplification of invertase alleles, and allele detection needed several PCRs. This phenomenon is known as allelic dropout and was observed in diverse studies in human genetics as well as in plant microsatellite genotyping (FINDLAY ET AL., 1995; TABERLET ET AL., 1996, BROQUET & PETIT, 2004; ZHANG ET AL., 2006; SOULSBURRY ET AL., 2007). Allelic dropout is the failure of PCR amplification of one allele in a heterozygous organism. Therefore, allele identification might be incomplete, and there is the possibility that not all existing invertase alleles of one genotype have been detected. Another aspect of allele mining by PCR was the definition of cDNA alleles, which were not detected as actively transcribed by pyrosequencing analysis of cDNA (section 3.2.2.1.1, Figure 3.2.24). A low number of cDNA clones represented these alleles and some occurred only once in PCR amplification. However, pyrosequencing analysis of genomic DNA showed that the SNPs specific for those alleles are present at genomic level, representing an existing allele of the genotype but is transcribed at such a low level, therefore not being detectable by cDNA pyrosequencing analysis. This is consistent with the rare amplification of these alleles by PCR.

Allele mining for each invertase gene was achieved by using tissues where invertase expression has been demonstrated (ZRENNER ET AL., 1996; MADDISON ET AL., 1999; HEDLEY ET AL., 1993, 1994). For the gene *invGF*, MADDISON ET AL. (1999) detected expression exclusively in floral tissues. Expression profiles of the tomato and the *Arabidopsis invGF* orthologs, *LIN7* and *ATβFRUCT2* respectively, showed also restriction of transcripts to floral tissues (FRIDMAN ET AL., 2003). In contrast to these findings, this study showed that *invGF* expression occurred also cultivar dependent in leaves (section 3.2.1.2, Figures 3.2.1 and 3.2.2). *invGF* alleles isolated from leaves and flowers were identical at amino acid level as well as in their nucleotide sequence.

It has to be pointed out that in the course of this study invertase allele mining using three tetraploid cultivars resulted in a tremendous allelic variation not expected by SSCP analysis that was earlier applied on 240 tetraploid potato cultivars (LI ET AL., 2005, 2008). Besides non synonymous allele specific SNPs, which caused amino acid exchanges, quite a number of synonymous SNPs occurred, leading to alleles identical at amino acid level. As synonymous SNPs do not cause amino acid substitutions, the function of the corresponding protein should not be affected. Nevertheless, several studies in humans showed that synonymous SNPs produced altered mRNA secondary structures affecting mRNA degradation and modification (e.g. splicing) as well as resulting in a reduced amount of translated protein (NACKLEY ET AL., 2006; reviewed in CHAMARY ET AL., 2006). Additionally, KIMCHI-SARFATY ET AL. (2007) showed that synonymous SNPs also affect enzymatic substrate specificities. In this respect, haplotypes showing synonymous SNPs accompanied by similar mRNA and protein levels, led to altered enzyme conformation because of rare codons, resulting from synonymous polymorphisms that affected the timing of co-translational folding and, therefore, the structure of the enzyme. Whether synonymous SNPs detected in potato invertase alleles also alter mRNA secondary structure, mRNA degradation, translation and enzyme conformation needs further investigation.

❖ Phenetic tree analysis of invertase cDNA alleles from all five potato invertase genes

The phenetic tree analysis was applied to group all alleles obtained from the five potato invertase genes according to their similarity at amino acid level (Figure 4.1). This allowed getting an impression of the grouping between the invertase isoforms as well as the clustering of associated and not associated alleles of the latter isoforms to each other.

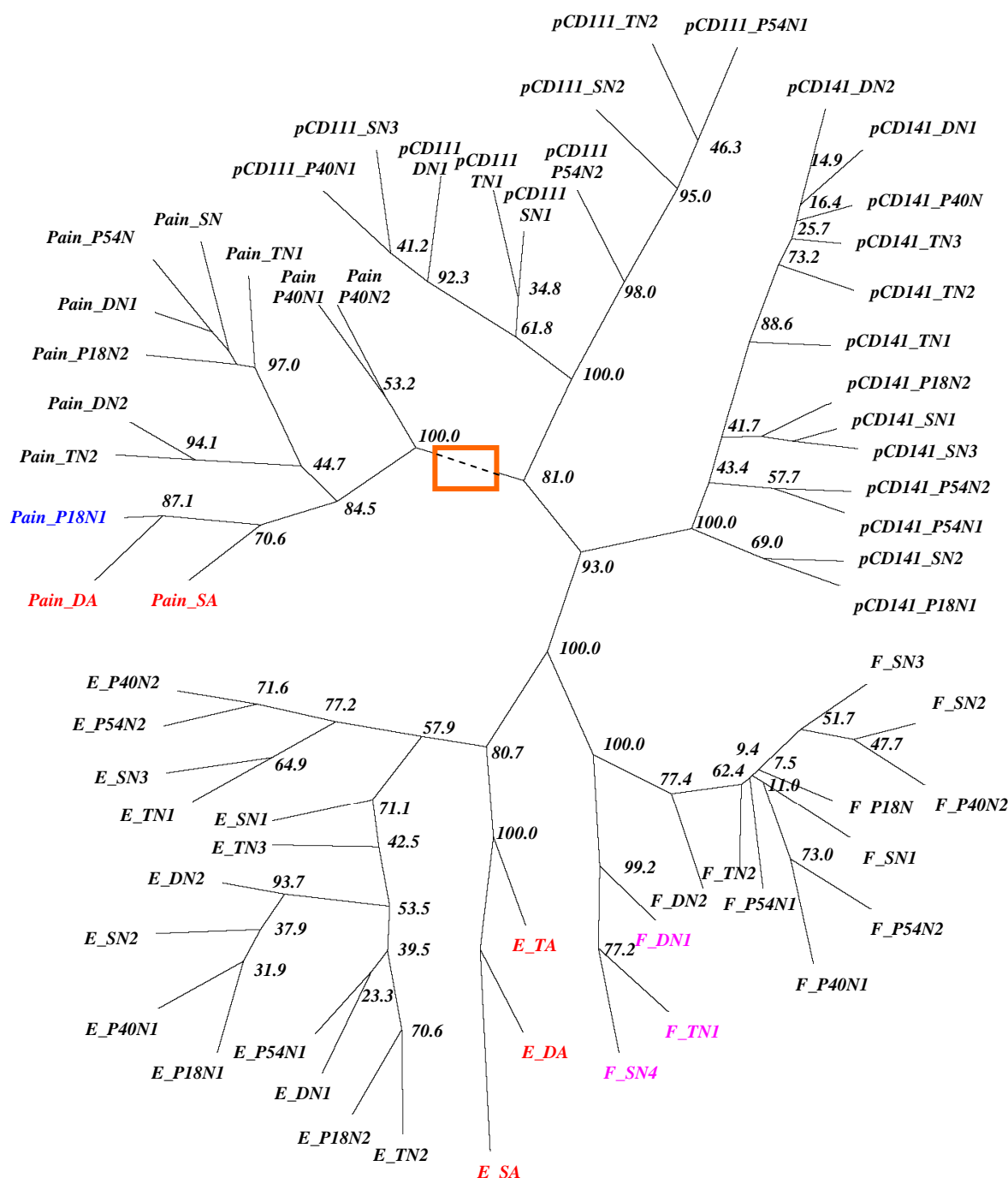


Figure 4.1: Amino acid based phenetic tree of all cloned invertase cDNA alleles. The tree was generated using the maximum parsimony method. A total of 100 bootstrapping runs were performed, and the percent reliability is indicated next to each branch. Allele names comprise the invertase gene, genotype and allele type. Alleles found to be associated with better potato chips quality are coloured in red. The allele *Pain_P18N1*, which is amino acid sequence identical to *Pain_DA* was not tested by association analysis and, therefore, is coloured in blue. The three *invGF* alleles *F_SN4*, *F_DN1*, and *F_TN1* are thought to be representatives of the associated SSCP fragment *invGF-4d* because of similar grouping as observed for the associated *invGE* alleles *E_SA*, *E_DA*, and *E_TA*. The three *invGF* alleles are coloured in pink. The boxed area reflects a truncated branch – this is due to the relatively distant clustering of vacuolar *Pain-1* alleles compared to cell wall-bound invertase isoforms (*invGE*, *invGF*, *pCD111*, and *pCD141*) because of weak sequence homology.

The phenetic tree of invertase alleles generated in this study revealed that the alleles from the different loci group separately. The analysis showed that the gene *Pain-1* is more similar to

the genes *pCD111* and *pCD141* because of closer branching distance in contrast to the genes *invGE* and *invGF*. Peptide based comparison of the different potato invertase isoforms revealed that the cell wall-bound invertases *invGE*, *invGF*, *pCD11,1* and *pCD141* share an amino acid sequence homology of approximately 73%, whilst the vacuolar invertase *Pain-1* displayed a homology of around 38% to the cell wall-bound isoforms explaining the distant clustering. This is in agreement with previous studies where comparison of amino acid sequences of plant invertases demonstrated that cell wall and vacuolar invertases belong to two different classes (TYMOWSKA-LALANNE & KREIS, 1998). Therefore, invertase amino acid sequences of the same isoform originating from different species are more similar to each other than sequences of different isoforms from one and the same species.

❖ Genomic characterization of the genes *Pain-1*, *pCD111*, and *pCD141*

The next step was to elucidate the genomic structures of the genes *Pain-1*, *pCD111*, and *pCD141*. Therefore, high density BAC library screens were performed using two different BAC libraries constructed with genomic DNA of the same diploid genotype (BALLVORA ET AL., 2002, 2007).

Information about genomic potato invertase organization was only available for the genes *invGE* and *invGF* on chromosome IX. MADDISON ET AL. (1999) showed that the *Inv_{ap-b}* locus consists of the genes *invGE* and *invGF* linked in a direct tandem repeat and separated by approximately 2.3kb from each other. Both genes exhibit a similar exon/intron structure composed of six exons and five introns. The size of the *Inv_{ap-b}* locus is approximately 8.6kb (MADDISON ET AL., 1999). Comparative analysis of the potato genetic map revealed that the genome segment harbouring the genes *pCD111* and *pCD141* on chromosome X is related to a genome segment containing the genes *invGE* and *invGF* of chromosome IX, and, therefore, might also be organized in a direct tandem repeat (GEBHARDT ET AL., 2003; FRIDMAN ET AL., 2003). Corresponding studies of orthologous invertase genes of tomato revealed that the syntenic tomato locus to *invGE/invGF*, *LIN5/LIN7* on chromosome IX, also displays the linkage of both genes in a direct tandem repeat (FRIDMAN ET AL., 2003). The tomato invertase genes *LIN8* and *LIN6* on tomato chromosome X, which are orthologs of the potato genes *pCD111* and *pCD141* on potato chromosome X, are also arranged in this manner. Known genomic organizations of above described potato and tomato invertase genes strongly encouraged the expectation that the potato genes *pCD111* and *pCD141* are also linked in a direct tandem repeat.

Sequencing of a BAC insert harbouring the genes *pCD111* and *pCD141* confirmed the latter spatial organization of both genes (section 3.3.1.3, Figures 3.3.11 and 3.3.12) as observed for

their tomato orthologs *LIN8* and *LIN6*. Furthermore, it could be shown that the genes *pCD111* and *pCD141* are separated from each other by approximately 8kb, whilst the tomato genes *LIN8* and *LIN6* are divided by 7,2kb. Additionally, the tomato locus *LIN8/LIN6* is flanked upstream by a kinesin, and downstream by a 40S ribosomal gene. These two genes were also detected flanking the potato gene pair *pCD111/pCD141* of the *Inv_{ap}-a* locus in the same orientation. The size of the *Inv_{ap}-a* locus is approximately 17.5kb.

The potato invertase genes *invGE* and *invGF* as well as the tomato invertase genes *LIN5*, *LIN7*, and *LIN6* consist of six exons and five introns. The tomato invertase gene *LIN8* displays five exons and four introns. However, genomic characterization of the potato genes *pCD111* and *pCD141* showed that both contain six exons and five introns.

The size of the exons and introns is not conserved between different potato invertase genes, except the nine base pair mini-exon II, and varies from 9 to 1,026 and from 83 to 2,682 nucleotides, respectively.

The gene *Pain-1* on chromosome III codes for a vacuolar invertase in contrast to *invGE/invGF* and *pCD111/pCD141*, which all encode cell wall-bound invertase isoforms. One interest of this study was whether *Pain-1* is differently organized than the other two known invertase loci. Searching for *Pain-1* tomato orthologs resulted in one EST on tomato chromosome III (www.sgn.cornell.edu). By sequencing BAC inserts harbouring the *Pain-1* gene no second invertase was detected. Within the BAC inserts *Pain-1* gene surrounding sequences should allow to detect a possible tandem repeat organization (section 3.1.1.4, Figure 3.1.11 A). This possibility was excluded for the *Pain-1* locus. Furthermore, in the corresponding flanking regions no kinesin or ribosomal genes as observed for the genes *pCD111/pCD141* were detected. The *Pain-1* gene consists of seven exons and six introns, and, therefore, has an additional exon and intron when compared to the other four known potato invertase genes. Analysis of invertase isoforms from different species like *Arabidopsis thaliana*, carrot, tomato, maize, tobacco, mung bean, and pea demonstrated that invertases vary in their exon/intron composition (TYMOWSKA-LALANNE & KREIS, 1998). All potato invertases contain the conserved exon II, which only consists of nine nucleotides. This mini-exon, one of the smallest exons known in plants, encodes the residues DPN of the highly conserved β -fructosidase motif, NDPNG (TYMOWSKA-LALANNE & KREIS, 1998). The β -fructosidase motif of the genes *invGE*, *invGF*, and *pCD111* is represented by NDPNA rather than NDPNG. The other potato invertases *Pain-1* and *pCD141* display the residues NDPNG. BOURNAY ET AL. (1996) observed that under cold stress the mini-exon of the *pCD111* cell

wall invertase was skipped in an alternative splicing event. The functional relevance of the splicing effect was not investigated.

Comparison of the amino acid sequences of plant invertases demonstrated that cell wall and vacuolar invertases belong to two different classes (TYMOWSKA-LALANNE & KREIS, 1998). Corresponding invertase amino acid sequences of the same isoform originating from different species are more similar to each other than sequences of different isoforms from one and the same species are. Additionally, typical vacuolar invertase peptide regions or residues were identified that are absent in cell wall-bound isoforms. The peptide domain WECxDF, which is conserved among plant invertases, where x=valine in vacuolar and x=proline in cell wall-bound invertases (TYMOWSKA-LALANNE & KREIS, 1998; STURM, 1999) was also present in the different invertase isoforms of potato.

❖ Structural 3D-analysis of invertase cDNA alleles

In collaboration with Pawel Durek (MPIMP/Golm) the structural consequences of allelic invertase sequence variation was explored by applying a 3D-modelling analysis. The modelling of the allelic invertase structure was based on the 3D crystal structure of cyanobacteria invertase (ALBERTO ET AL., 2004). The models presented included the putative sucrose binding domain. In addition to the structural visualization of amino acid exchanges, also the electric potential (EP) of *Pain-I* and *invGE/invGF* alleles was mapped at pH 4.7 mimicking the vacuolar and apoplastic environment.

The analysis of the molecular structure of the associated ‘Satina’ *Pain-I* allele *Pain_SA* compared to *Pain_SN3* and of the associated ‘Diana’ allele *Pain_DA* compared to *Pain_DN1* and *Pain_DN2* showed that amino acid exchanges were manifested on the surface of the enzyme. None of the modelled molecules showed structural differences within the putative sucrose binding domain (section 3.1.1.3.1, Figure 3.1.7).

Comparative models of ‘Diana’ alleles revealed that models of the two not associated alleles *Pain_DN1* and *Pain_DN2* differed less from each other compared to the associated allele (*Pain_DA* vs. *Pain_DN1*; *Pain_DA* vs. *Pain_DN2*). The SNP 1544 present in the alleles *Pain_SA* and *Pain_DA*, which was found to be associated with better potato chips quality, had a direct effect on the molecule’s surface.

The characterization of the EP showed charge differences among the protein models. The putative sucrose binding site was positively charged matching the partial negative charge of the substrate sucrose due to the hydroxyl groups. A dramatic charge difference of the putative sucrose binding site was observed in the associated *Pain-I* alleles *Pain_SA* and *Pain_DA*, which are not amino acid identical but share the associated SNP at cDNA position 1544

(section 3.1.1.3.2, Figure 3.1.9). The EP switched from positive to neutral compared to the not associated *Pain-1* alleles. The EP changes could not be correlated to amino acid exchanges near the sucrose binding site. The causative amino acids were not yet analyzed, but are subject of ongoing investigations in the research project. Whether these EP differences of the associated alleles compared to the EP of the not associated alleles might influence enzymatic activity remains unclear. Possible effects of the neutral charge might be a weaker binding of sucrose and, therefore, a reduced conversion of sucrose into the reducing sugars glucose and fructose, which influence negatively potato chips quality. This would support the hypothesis that sequence variation of potato invertases leads to functional variation and that in consequence associated alleles produce less reducing sugars in tubers and thus a good potato chips quality.

Modelling *invGE* alleles from the cultivars ‘Satina’ and ‘Theresa’ revealed similar molecular structures of the investigated alleles (section 3.2.1.3, Figure 3.2.14). Only slight differences were visible. Comparative modelling of the associated *invGE* alleles *E_SA* and *E_TA* did not reveal strong differences (Figure 3.2.15). Superimposing the associated *invGE* allele *E_SA* and the *invGF* allele *F_SN4* showed strong structural differences (Figure 3.2.23). The strong structural effects are likely caused by sequence divergence of the two genes. Even though the latter belong to the same locus and arose from gene duplication (GEBHARDT ET AL., 2003; FRIDMAN ET AL., 2003), *invGE* and *invGF* proteins are only 74% similar to each other (MADDISON ET AL., 1999).

EP analysis of *invGE* alleles (Figure 3.2.16 and 3.2.17) revealed only weak charge differences of the associated ‘Satina’ allele *E_SA*, which showed a small negatively charged area compared to the allele *E_SN3*. The associated ‘Theresa’ allele *E_TA* which differs in its amino acid sequences to *E_SA* but contains the associated SNP 1103*, exhibited no charge differences of the putative sucrose binding site compared to *E_TN1*.

For the gene *invGF* two alleles of the cultivar ‘Satina’ *F_SN3* and *F_SN4* were modelled (section 3.2.1.3, Figure 3.2.19). Comparison of both allelic structures showed differences among the molecules. EP analysis showed that especially the putative sucrose binding domain displayed visible changes of its charge between the two alleles. In the allele *F_SN4* the EP of the putative domain switched from positive to negative, influencing possible interactions of substrate and enzyme (Figure 3.2.21). As reported for the *Pain-1* and *invGE* alleles, EP changes are not directly mediated by amino acid exchanges nearby the binding site. It is most likely that surrounding amino acid exchanges cause the charge shift.

In conclusion, 3D-modelling is a tool to gain first insights in the possible structural consequences of different invertase alleles. The models nicely showed that amino acid exchanges can modify the surface of the enzyme. However, no effect on the structure of the putative sucrose binding domain was detected in any of the modelled molecules, but an effect on the charge was observed. The putative sucrose binding site exposed areas of neutral and positive charges in the associated *Pain-1* and *invGE* alleles, *Pain-SA*, *Pain_DA*, and *E_SA*, and in the putative associated *invGF* allele *F_SN4*. Whether these EP differences of the alleles influence enzymatic activity remains unclear. To address this question a biochemical characterization was carried out (*Pain-1*: section 3.1.2.3; *invGE/invGF*: section 3.2.2.3) to determine whether differences in enzymatic activity are present between invertase alleles. The corresponding results will be discussed in one of the following section (4.3: Biochemical analysis of *Pain-1*, *invGE*, and *invGF* alleles).

Extracellular invertase from yeast and alkaline invertase from *Vicia faba* are known to act as oligomers (KERN ET AL., 1992; ROSS ET AL., 1996). Due to 3D-analysis, it might be possible that the vacuolar and cell wall-bound potato invertases might also function as a higher complex of several monomers. ROSS ET AL. (1996) did not investigate the role of allelic subunits in the alkaline invertase complex. Supported by putative 3D structural analysis, there is the possibility that invertase enzyme complexes are formed by allelic proteins (Pawel Durek, personal communication). Considering this information, it is comprehensive that conformational and electrostatic changes on the enzyme's surface affect invertase enzyme complex formation and might lead to altered enzyme activity as reported for other studies about complex formation of enzymes (TETLOW ET AL., 2004a; TETLOW ET AL., 2004b; TETLOW ET AL., 2008).

4.3 Functional characterization of potato invertase alleles

The expression of single genotype specific *Pain-1*, *invGE*, and *invGF* alleles was quantified with pyrosequencing using cDNA. Allele specific transcript abundance was compared to the genomic allele dosages as determined by pyrosequencing using genomic DNA. Additionally, total amounts of *Pain-1* transcripts during tuber cold storage were determined using qRT-PCR.

Pyrosequencing is a robust and quantitative sequencing method, based on real-time detection of pyrophosphate, which is released as a result of nucleotide incorporation in a sequencing-by-synthesis reaction (RONAGHI ET AL., 1996). Assessing allele frequencies in large genomic DNA pools by pyrosequencing has demonstrated the high level of accuracy of this method. WASSON ET AL. (2002) reported the reliable detection of allele frequency differences of 4% between DNA pools from human populations. This result was confirmed by NEVE ET AL. (2002) who estimated that, for large DNA pools, allele frequencies that differ by $\pm 5.2\%$ would be significant. Because of its high accuracy, pyrosequencing has been considered as the method of choice for genotyping SNPs in polyploid species. For example, RICKERT ET AL. (2002) and OEFNER (2002) have shown that the different heterozygous states of a binary SNP in tetraploid potato could reliably be distinguished. Furthermore, RICKERT ET AL. (2002) showed that 82% of the polymorphic sites tested were amenable to allelic discrimination by pyrosequencing, which is by far better than any other SNP genotyping method. Another advantage of pyrosequencing is the possibility to determine multiple SNP frequencies in a single measurement, allowing analysis of more than two alleles simultaneously.

The biochemical analysis of invertase cDNA alleles was performed using yeast as heterologous expression system. The complementation of a yeast invertase mutant with potato cDNA alleles allowed expression of single alleles and their characterization. Invertase activities were assayed to determine substrate affinities (K_m) and reaction rates (v_{max}).

The genes *pCD111* and *pCD141* were not functionally analyzed because of limited information concerning association and putative allelic effects on potato chips quality at the beginning of this work.

In a simple scenario, one can imagine several functional characteristics of an invertase allele influencing positively potato chips quality. Even though potato chips quality is a multigenic trait, invertase was the first gene studied at functional level in terms of allelic differences contributing to this trait. Possible features of a superior allele are summarized in Figure 4.2.

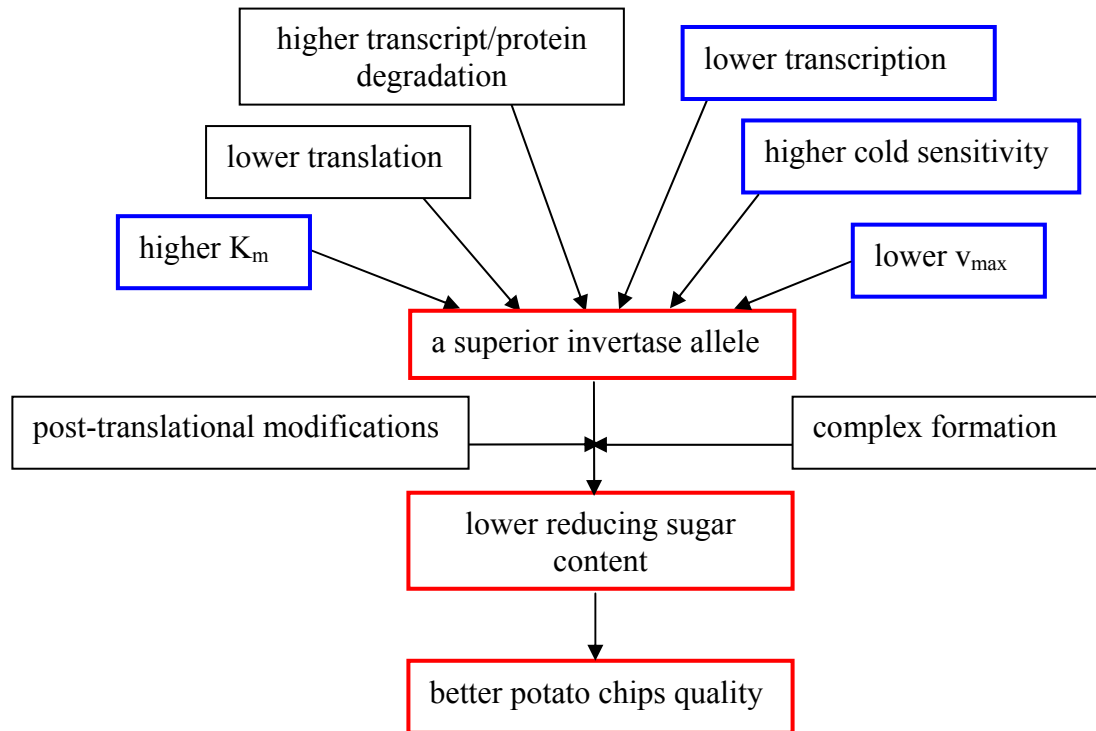


Figure 4.2: Simplified scheme displaying features of an ‘ideal’ invertase allele acting positively on potato chips quality. Red boxes indicate desired phenotypic traits influenced by a superior invertase allele. Blue boxes represent functional characteristics that were studied during this work.

A superior invertase allele should lead to a low reducing sugar content, which in consequence gives rise to a good potato chips quality. Allelic effects can occur at transcriptional, translational, post-translational, and biochemical level. In this agreement, a superior invertase allele would be sparsely transcribed and translated, displaying low protein abundance, and, therefore, having a poor contribution to overall invertase activity consequently leading to a reduced sucrose conversion. Additionally, high levels of transcript and protein degradation would also contribute to low protein content. In addition, a superior invertase allele could be characterized by a high K_m value meaning a low affinity to sucrose combined with a low v_{max} value indicating a slow conversion of sucrose. These biochemical characteristics can be even more prevalent in a cold environment applied to potato tubers to prevent them from sprouting. However, other post-translational modifications like N-glycosylation, protein folding, or allelic complex formation are further possibilities in influencing a superior invertase allele resulting in lower reducing sugar content and, therefore, better potato chips quality.

In the course of this work, out of the mentioned features characterizing a superior invertase allele, allele specific expression and allelic biochemical characteristics were determined to prove whether such alleles are present.

❖ Expression analysis of *Pain-1*, *invGE*, and *invGF* alleles

Besides the possibility of an altered enzymatic activity, sequence variation can also manifest at expression level. A simple assumption consist of a low transcription level of a superior invertase allele resulting in low translation rate contributing to a lower extent to overall invertase activity due to limited enzyme abundance. In the latter case lower invertase protein levels due to a reduced allelic expression would lead to superior chips quality (Figure 4.2).

Differential expression analysis was performed monitoring the genes *Pain-1*, *invGE*, and *invGF*. The study of the expression followed different purposes. The gene *Pain-1* is known to be differentially expressed during tuber cold storage (ZRENNER ET AL., 1996; ZHOU ET AL., 2004; BAGNARESI ET AL., 2008). Using pyrosequencing, genotype specific allelic transcription patterns during a cold storage time course of four weeks were determined compared to additional measured genomic dosages of the alleles. Total amounts of *Pain-1* transcripts during tuber cold storage were measured using qRT-PCR.

The expression of *invGE* and *invGF* alleles was detected by pyrosequencing analysis in comparison to the allelic genomic dosage determined by pyrosequencing using genomic DNA as template.

Expression analysis of *Pain-1*, *invGE*, and *invGF* alleles revealed differential transcriptional regulations of particular alleles. Allele specific expression is a common fact and several studies showed such a regulation on transcriptional level (reviewed in KNIGHT, 2004). Studies in octoploid strawberry showed allele specific expression of a pathogenesis-related gene induced by fungal infection causing fruit rot (SCHAART ET AL., 2005). SPRINGER & STUPAR (2007) used allele specific expression assays to profile the relative allelic expression in seedling tissue derived from maize hybrids. They found evidence for regulatory variation that contributes to biased allelic expression between genotypes and between tissues. Studies in mammals during early stages of development demonstrated the importance of allele specific expression implicating imprinting mechanisms (SZABO & MANN, 1995).

The alleles *Pain_SA* and *Pain_DA* were derived from the genotypes ‘Satina’ and ‘Diana’. It could be shown that the two alleles, having different amino acid sequences but being associated with better potato chips quality, were differentially expressed during tuber cold storage when compared to their genomic dosage (section 3.1.2.1, Figures 3.1.12 and 3.1.14). Whilst *Pain_SA* showed an up-regulation in its expression of approximately 18%, the allele *Pain-DA* revealed a permanent reduction of expression during the cold storage course of four weeks resulting in an overall decrease up to 20% compared to samples that were not stored in the cold. The diploid potato genotype P18 harbours an allele with identical amino acid

sequence to the associated allele *Pain_DA*. The allele *Pain_P18N1* showed a strong transcript increase after one week of tuber cold storage (section 3.1.2.1, Figure 3.1.18). Expression level was 50% higher than in tubers that were not stored in the cold. The differential expression of the associated allele *Pain_SA* and the allele *Pain_P18N1*, which is identical at amino acid level to the associated allele *Pain_DA*, showed that these allelic transcripts are influenced by low temperature leading to higher abundance during tuber cold storage than expected from their genomic dosage. This observation is in contradiction with the hypothesis that a superior invertase allele is characterized by transcriptional regulation implicating a lower expression (Figure 4.2). The associated allele *Pain_DA* exhibited a lower transcription level during tuber cold storage showing the lowest abundance after four weeks in the cold. This is in agreement with the assumption mentioned above that superior alleles display low amounts of transcripts, protein and, therefore, a lower enzymatic activity. However, the impact of the ‘Diana’ allele on overall invertase activity remains unclear since no methods are available to measure the activity of single allelic invertases *in vivo*. Additionally, tissue or clonal specific enzyme activity due to transcriptional changes in specific tissues or single cells might contribute to altered expression patterns and, therefore, lead to a quite variable enzyme activity.

The other alleles not associated with superior chips quality of the genotypes ‘Diana’, ‘Theresa’, and P40 revealed no relevant changes in their expression pattern during tuber cold storage. The allelic expression remained similar compared to the genomic dosage of the alleles.

Allelic differences and the contrasting observations regarding expression of the associated alleles at transcript level might be due to the individual genetic background. Genotype specific transcription factors and other regulatory elements as well as allele specific promoter sequences can result in differences of allele expression. Additionally, observations of allele specific expression patterns are limited due to the limited number of analyzable genotypes in the course of this work.

Using qRT-PCR, total *Pain-I* transcripts were quantified during tuber cold storage. ZRENNER ET AL. (1996) detected soluble acid invertase transcripts after 44h of tuber cold treatment at 4°C using Northern blots. Transcript levels reached a maximum after seven days of cold storage, followed by a decrease throughout the next two to six weeks. In contrast, BAGNARESI ET AL. (2008) detected an increase of soluble acid invertase transcripts at week two to four of cold storage.

In this study expression analysis showed an intense up-regulation of *Pain-I* transcripts after one to two weeks of tuber cold storage at 4°C. After the first week of cold treatment P18

transcripts showed the highest level of relative expression compared to all other analyzed genotypes (section 3.1.2.1, Figure 3.1.19). The cultivar ‘Theresa’ was the only genotype where total invertase transcripts increased again after four weeks of tuber cold storage (section 3.1.2.1, Figure 3.1.17).

The results of the expression analysis of the soluble acid invertase *Pain-1* are in agreement with earlier observations by ZRENNER ET AL. (1996) and BAGNARESI ET AL. (2008). Additionally, it was demonstrated that in different genotypes *Pain-1* alleles were strongly cold influenced regarding their expression, whilst others showed minor changes or stayed unaffected compared to their genomic distribution (section 3.1.2.1, Figures 3.1.14 and 3.1.16).

Pyrosequencing of *invGE* and *invGF* alleles of leaf and floral tissue respectively, showed that genotype specific alleles are differentially expressed compared to their genomic dosage. The question might arise how alleles analyzed in leaves and flowers might contribute to chips quality when not acting directly in the potato tuber. Association analysis of *invGE* and *invGF* genes revealed positive correlations between corresponding allelic fragments and potato chips quality (LI ET AL., 2005). Expression profiles investigated by MADDISON ET AL. (1999) using GUS histochemical expression analysis showed that *invGE* is expressed in several tissues and also under the ‘eyes’ of the tuber. In this study, none of the six selected genotypes allowed *invGE* transcript amplification using cDNA synthesized from mature tuber ‘eye’ RNA. Therefore, leaf tissue has been chosen to identify *invGE* alleles as well as to study their expression. It might be possible that mature tubers were not the appropriate source for *invGE* allele mining even though MADDISON ET AL. (1999) also used mature tubers but from a different cultivar. In this study mature tubers were used to extract RNA. Following the study’s approach analyzing potato chips quality before and after tuber cold storage, tubers were directly processed after harvest or stored in the cold for one to four weeks. MADDISON ET AL. (1999) also showed that *invGF* expression is restricted to floral tissues in the cultivars ‘Désirée’ and ‘Saturna’, which was shown to be not consistent with findings concerning some of the selected genotypes in this study where *invGF* expression was also detected in leaves (section 3.2.1.2, Figures 3.2.1, 3.2.2).

Reasons for the lack of *invGE* transcripts in potato tubers might also be that under the latter conditions *invGE* expression is below the detection limit due to time-dependent expression pattern. Cell wall-bound invertases are known to play an important role in tuber initiation. MINHAS & SAINI (2004/5) showed that under tuber-inducing conditions cell wall invertase activity increases up to 74% during the stolon to tuber transition period. Cell wall-bound invertase has been recognized as a key enzyme in apoplastic phloem unloading, which

switches in the developing tuber to the symplastic mode resulting in the primary function of sucrose synthase (SuSy) in sucrose cleavage. Cell wall invertases regulate the import of the reducing sugars glucose and fructose, which are used for starch synthesis in stolon tips. Assuming a superior invertase allele, less transcribed and translated resulting in low protein abundance accompanied by low activity, might lead to retarded sucrose cleavage and, therefore, less import of sugars in the initiated tuber. Additionally, a less active invertase allele in the apoplast might influence sink strength in a way that phloem unloading is decelerated and further tuber storage compounds are less abundant. Besides playing an important role in tuber initiation, cell wall invertases are generally believed as main determinants of sink strength especially during the initial stages of sink development (ROITSCH, 1999; ROITSCH ET AL., 2000). Various experimental approaches have been used to demonstrate the importance of cell wall-bound invertase isoforms for assimilate partitioning and determining sink strength. These include inhibition of storage tissue development in carrot roots by antisense repression of cell wall invertase (STURM & TANG, 1999; TANG ET AL., 1999), increase in potato tuber size by over-expression of cell wall invertase (TAUBENBERGER ET AL., 1999), arrested seed development in maize mutant lacking cell wall invertase (MILLER & CHOUREY, 1992), induction of sink metabolism in source leaves of transgenic plants by over-expression of a yeast invertase (STITT & SONNEWALD, 1995), and specific expression of a cell wall invertase during pre-storage phase in the thin walled parenchyma of faba bean seed coat (WEBER ET AL., 1995).

A possible explanation for the association of *invGF* alleles transcribed in flowers and leaves might be the fact of LD with associated *invGE* alleles detected in SCCP based analysis (LI ET AL., 2005). It might be comprehensive to think of a haplotype block associated with better chips quality including both genes *invGE* and *invGF*.

The associated *invGE* alleles correlated with better potato chips quality (LI ET AL., 2005) revealed minor changes in their expression in leaves compared to their genomic dosage (section 3.2.2.2.1, Figure 3.2.24). The ‘Satina’ allele *E_SA* and the ‘Diana’ allele *E_DA*, which are identical in their nucleotide and amino acid sequences, are present in simplex (25%) in the corresponding genotypes. Allele expression was reduced by 5% to a level of 20% in leaves. The associated ‘Theresa’ allele *E_TA*, which is present in simplex (25%) and is different in its amino acid sequence to *E_SA* and *E_DA*, showed a slight expression induction up to 35% in leaves. Transcripts for the ‘Diana’ allele *E_DN2* and the ‘Theresa’ allele *E_TN3* were not detected in pyrosequencing analysis possibly due to transcripts below the detection level. These findings indicated an allele specific expression of invertase genes in

potato. Whether this differential expression of *invGE* alleles influences potato chips quality remains unclear since no transcripts in tubers were detectable.

The *invGE* alleles of the diploid potato genotypes P18, P40, and P54 were differentially expressed in leaves as expected from their genomic dosage (section 3.2.2.1.2, Figure 3.2.25). Transcripts of the alleles *E_P18N1*, *E_P40N2*, and *E_P54N1* were more abundant, whilst the other alleles were less present compared to the genomic distribution.

Pyrosequencing based expression analysis of *invGF* alleles showed that the alleles *F_SN4*, *F_DN1*, and *F_TN1*, which group separately from the other *invGF* alleles were not expressed differentially in flowers compared to their genomic dosage (section 3.2.2.1.3, Figure 3.2.26). All three alleles are present in simplex (25%) in the corresponding genotypes. The ‘Satina’ allele *F_SN4* and the ‘Theresa’ allele *F_TN1* are identical at nucleotide and amino acid level. It is speculated that *F_SN4*, *F_DN1*, and *F_TN1* might refer to the *invGF-4d* SSCP fragment, which was found to be associated with better potato chips quality. The speculation is based on the fact that the three alleles show the same separation like the associated *invGE* alleles *E_SA*, *E_DA*, and *E_TA* in phenetic tree analysis (*invGE*: section 3.2.1.2.4, Figure 3.2.6; *invGF*: section 3.2.1.2.8, Figure 3.2.11). Additionally, the ‘Satina’ allele *F_SN3* showed a strong decrease in transcripts, whilst *F_SN1* was the prevalent allele in flowers compared to the allelic genomic distribution. The allocation of the alleles from the genotypes ‘Diana’, ‘Theresa’, and P40 did not differ. The two alleles of P54 revealed minor changes, the allele *F_P54N1* was decreased, whilst the allele *F_P54N2* was increased compared to their genomic distribution.

In conclusion, expression pattern of *Pain-I*, *invGE*, and *invGF* alleles of six genotypes did not correlate with association analysis, which was based on 240 tetraploid individuals (LI ET AL., 2005, 2008). The observation of genotype and invertase gene dependent transcriptional changes might be due to the genetic background. Since no allelic promoters or transcription factors and other modifiers were analyzed in this study the impact of these on allelic invertase expression cannot be clearly assessed. Nevertheless, the results showed that invertase genes are expressed in a genotype specific manner and that associated alleles followed divergent expression patterns. As expression QTLs (eQTLs) are an emerging field of interest in plants such analysis will be an interesting extension of the results gained in the present study, although expression data could not directly be linked to protein abundance and/or enzymatic activity.

❖ The heterologous system yeast

Aiming at the biochemical characterization of single invertase alleles, the use of a heterologous system was necessary to separate alleles from each other. With the knowledge that plant invertases are functional in yeast (FRIDMAN ET AL., 2004), the yeast invertase mutant *SUC2* (GOZALBO & HOHMANN, 1989), which lacks invertase activity was chosen for potato invertase allele expression. FRIDMAN ET AL. (2004) demonstrated the functional complementation of a yeast mutant with tomato cell wall-bound invertases. In this study it was shown that also a vacuolar invertase isoform complements the yeast *SUC2* mutant phenotype. All *SUC2* transformants harbouring *Pain-1*, *invGE*, and *invGF* cDNA alleles were able to grow on sucrose as sole carbohydrate source indicating functional complementation. The cDNA alleles of all three different genes were expressed under the control of the constitutive promoter *Adh1* to obtain high amounts of potato invertase protein.

With respect to the yeast and potato codon usages, it was found that the translation efficiency between both systems is different leading to less potato invertase protein depending on the allelic nucleotide sequence. It was tried to balance this translation deficiencies by incubation of complemented yeast strains for three days achieving equal amounts of potato invertase protein. Immunoblot analysis of *Pain-1* yeast transformants reflected the equal distribution of allelic invertase proteins (section 3.1.2.4, Figures 3.1.24 and 3.1.25).

In yeast, two invertase isoforms occur, which are encoded by the same gene, but originate from differential splicing events. One isoform is active, N-glycosylated, and extracellular, the other isoform is inactive, nonglycosylated, and located in the cytoplasm. The extracellular yeast invertase is targeted for secretion by a signal peptide, which is then removed by a peptidase. Plant vacuolar and cell wall-bound invertases are synthesized as prepropeptides (TYMOWSKA-LALANNE & KREIS, 1998). These peptides consist of N-terminal extensions up to 100 amino acid residues in length, which harbour a signal peptide and an N-terminal propeptide (STURM, 1999). The signal peptide is required for the entry in the endoplasmatic reticulum (ER), which leads to the secretory pathway. The N-terminal propeptide as well as a C-terminal extension, the latter being characteristic for vacuolar invertases, are thought to act as vacuolar sorting signals (MATSUOKA & NAKAMURA, 1991). Figure 4.3 illustrates vacuolar and cell wall-bound preproproteins and included domains (adopted from TYMOWSKA-LALANNE & KREIS, 1998; STURM, 1999).

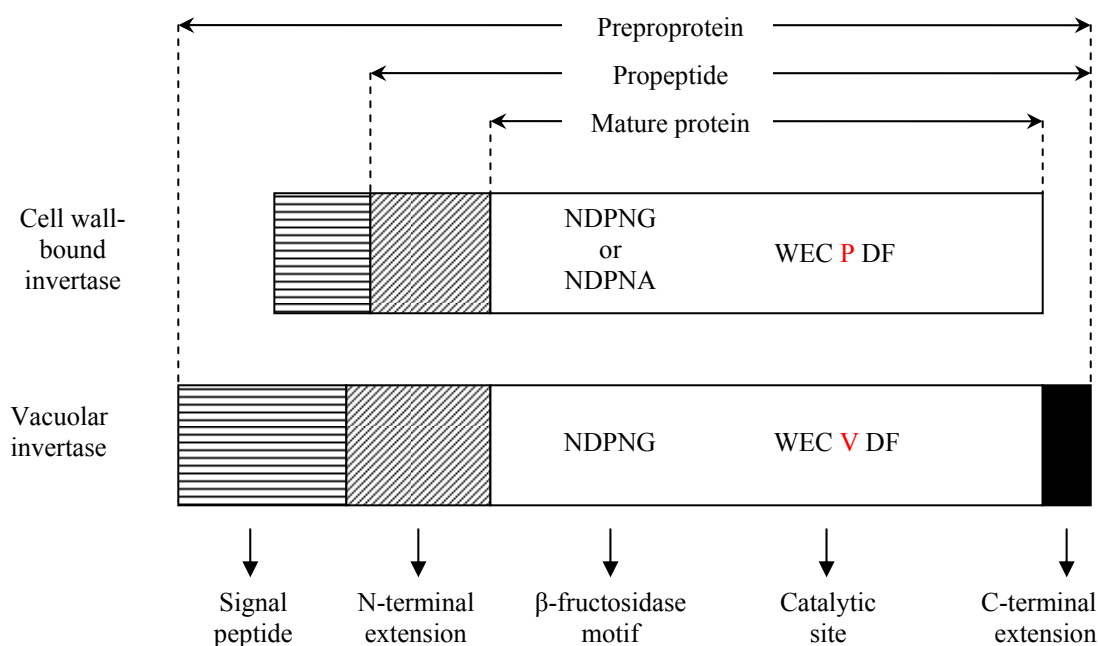


Figure 4.3: Schematic comparison of vacuolar and cell wall-bound preproprotein invertases. The preproprotein sequence includes the signal peptide, N- and C-terminal extensions and the mature protein. The peptide sequences NDPNG/NDPNA and WEC^{P/V}DF represent the β -fructosidase motif and the catalytic site, respectively.

Since yeast has no vacuole or a similar compartment, the question arises how vacuolar allelic potato invertases are processed and targeted in yeast to remain functional. This question can not be answered although functional complementation of the yeast invertase mutant was achieved.

Yeast invertase is known to be an oligomeric glycoprotein with 14 potential N-glycosylation sites located in the sequence (REDDY ET AL., 1999). The carbohydrate chains play a role in structure, function, stability, and folding of glycoproteins (KERN ET AL., 1992). In the yeast Golgi apparatus, N-glycosylation is achieved by a varying number of extended outer polymannose chains, yielding a high-mannose-type glycosylated protein with an average of nine to ten oligosaccharides per peptide chain. The glycosylated proteins are strongly heterogeneous in their carbohydrate content. Studies with yeast invertase demonstrated that glycosylation is a prerequisite for the oligomerization of the enzyme beyond the state of the active dimer. It was shown that core-glycosylation of yeast invertase is necessary for tetramer and octamer formation (KERN ET AL., 1992).

In plants, vacuolar and cell wall-bound invertases are glycosylated, whilst cytoplasmatic invertase isoforms are not glycosylated (STOMMEL & SIMON, 1990). Glycosylation of vacuolar and extracellular glycoproteins are different from each other. Most of the vacuolar glycoproteins described so far were found to be N-glycosylated with modified N-glycans containing fucose and/or xylose residues, but devoid of terminal glucosamine residues. In

contrast, extracellular glycoproteins were found to be N-glycosylated mostly by complex-type N-glycans including large structures with terminal fucose and galactose residues (RAYON ET AL., 1998). Both types of glycoproteins are post-Golgi modified leading to protein maturation in the vacuole or in the extracellular compartment. It was shown that N-glycosylation in plants plays an important role in prevention of proteolytic degradation, induction of correct folding, and biological activity of the protein. Furthermore, N-linked oligosaccharides may contain targeting information, or may be involved in protein recognition (RAYON ET AL., 1998). FAYE & CHRISPEELS (1989) found that unglycosylated cell wall-bound invertase of carrot was degraded in the secretory pathway or immediately after attaining the cell wall. As indicated above, N-glycosylation of proteins in plants is more complex than in yeast. It might be possible that the analyzed potato invertase alleles were not modified according their original status in the heterologous system yeast. Following the argumentation that N-glycosylation plays an important role in correct protein folding, yeast modified potato alleles might not be able to fold correctly. Due to possible folding deficiencies and less complex N-glycosylation, which both is crucial for enzyme activity, it might be also reasonable that measured potato invertase activity does not reflect the native situation in potato. Additionally, the invertase oligomerization must be considered. Yeast invertase was found to be a mixture of dimers, tetramers, and octamers (KERN ET AL., 1992). Studies in *Vicia faba* on alkaline invertase assumed that this enzyme acts as a homotetramer (ROSS ET AL., 1996). Considering this information, it might be possible that soluble acid invertases as well as cell wall-bound invertase isoforms might also function in a complex of subunits. In none of the mentioned oligomeric analysis, the role of allelic monomers was investigated. Assuming that a functional potato invertase complex is build of different allelic subunits in different ratios, yeast transformants containing only one allele do not allow the assembly of ‘multi-allelic’ complexes. VEITIA ET AL. (2008) reviewed cellular reactions to gene dosage imbalances and suppose that stoichiometric imbalances in macromolecular complexes are source of dosage-dependent phenotypes. In this context, it might be reasonable that putative invertase complexes being ‘mono-allelic’ in yeast do not reflect the *in vivo* situation in potato.

❖ Biochemical analysis of *Pain-1*, *invGE*, and *invGF* alleles

Biochemical analysis of plant invertases in heterologous systems was successfully demonstrated for a tomato cell wall-bound invertase isoform by FRIDMAN ET AL. (2004). In that study an invertase allele originating from a wild species (*S. pennellii*), which is barely functional indicated by low fruit sugar content was compared to the homologous invertase allele of the cultivated tomato (*S. lycopersicum*). In contrast, the present study aimed towards

the characterization of functional potato invertase alleles from different genotypes to detect allelic enzyme activities. The possible functional variability of invertases operates under field conditions and, therefore, should not have severe effects on fitness and might be difficult to detect.

Using yeast, the challenge was to determine possible minor enzymatic allelic differences.

The heterologous system yeast made it possible to measure enzyme activity of single potato invertase alleles. The enzymatic parameters Michaelis constant (K_m) and maximal velocity (v_{max}) of the invertase reaction were determined.

The biochemical analysis was carried out for 14 *Pain-I*, 10 *invGE*, and 11 *invGF* invertase alleles. For alleles of the genes *invGE* and *invGF* of the *Invap-b* locus only K_m values are presented because of missing a suitable antibody for determination of the invertase protein levels to evaluate v_{max} values, which are enzyme dependent.

Considering ideal biochemical characteristics of a superior invertase allele, a high K_m value demonstrating the low substrate affinity of the enzyme accompanied by a low v_{max} value representing a slow conversion of sucrose would be a basic working model (Figure 4.2). Additionally, a higher cold sensitivity of the enzyme can intensify the described biochemical parameters leading to even less sucrose conversion into the reducing sugars glucose and fructose and a better potato chips quality.

Biochemical analysis of the *Pain-I* alleles was performed at 30°C and at 4°C. The analysis at 4°C was carried out to study possible differences of enzyme kinetics due to allelic amino acid composition in response to cold storage conditions. It was previously reported that cold storage influences transcriptional changes regarding vacuolar invertases (ZRENNER ET AL., 1996; ZHOU ET AL., 2004; BAGNARESI ET AL., 2008), but no study investigated the effect of low temperatures on the enzyme activity itself. The 30°C assay of *Pain-I* alleles showed that substrate affinity ranged between 15mM of the allele *Pain_P40N2* and 23mM of the allele *Pain_SN* (section 3.1.2.3, Table 3.1.15). Maximal velocities of the alleles varied between 2mmol*h⁻¹*mg protein⁻¹ in the case of the allele *Pain_TN2* and 12mmol*h⁻¹*mg protein⁻¹ for the allele *Pain_DA*. The enzymatic characteristics of the analyzed alleles from the cultivar ‘Satina’ showed no significant differences. In the K_m and v_{max} values of the cultivar ‘Diana’ differences were displayed in respect to the allele *Pain_DN2*. *Pain_DN2* showed the highest substrate affinity with a K_m of approximately 16mM, and the slowest substrate conversion with a v_{max} of around 6mmol*h⁻¹*mg protein⁻¹. The other two analyzed alleles *Pain_DA* and *Pain_DN1* displayed substrate affinities of approximately 20mM and maximal velocities of around 12mmol*h⁻¹*mg protein⁻¹. Comparison of the associated *Pain-I* alleles *Pain_SA* and

Pain_DA, which are not identical at amino acid level, revealed no significant differences regarding the enzyme's affinity to sucrose, but showed strong differences in the rate of sucrose conversion. The allele *Pain_DA* converts sucrose approximately 2.5 times faster than the allele *Pain_SA*.

Looking at the other analyzed *Pain-I* alleles, the two 'Theresa' alleles showed similar enzymatic characteristics and did not differ significantly. K_m values varied from 20 to 21mM and v_{max} values ranged from 2 to 3mmol*h⁻¹*mg protein⁻¹. Biochemical analysis of the alleles from the diploid potato genotypes P18 and P40 did not display any significant K_m and v_{max} differences. K_m - and v_{max} values of *Pain_P18N1* and *Pain_P18N2* ranged from 17 to 20mM and 3 to 5.5mmol*h⁻¹*mg protein⁻¹, respectively. The alleles *Pain_P40N1* and *Pain_P40N2* displayed substrate affinities between 15 and 17mM and v_{max} values between 6 and 7mmol*h⁻¹*mg protein⁻¹.

The biochemical analysis of the enzymatic characteristics of the *Pain-I* alleles at 4°C showed a dramatic increase in the enzyme's affinity to sucrose (section 3.1.2.3, Table 3.1.15). It is accepted that the rate of chemical reaction decreases approximately twofold for each 10°C decrease in temperature (AVERY, 1974). The K_m values decreased approximately 5.5 times compared to K_m values measured at 30°C. Also the maximal velocities of the alleles were affected at 4°C. By trend v_{max} values were around 2mmol*h⁻¹*mg protein⁻¹. The 4°C invertase assay showed the effect of low temperatures towards the vacuolar invertase alleles of potato. In the cold, the alleles showed a higher affinity to sucrose and in agreement a slower maximal velocity due to a stronger binding of the substrate and the general effect of low temperature to slow down enzymatic reaction rates. In contrast, the sucrose affinity of the yeast wild type strain *FY 1679*, used as reference, was not that much affected in the cold. The K_m values remained stable at approximately 22mM. Differences were detected in the rate of sucrose conversion. Yeast invertase slowed down from approximately 24mmol*h⁻¹*mg protein⁻¹ to 7.5mmol*h⁻¹*mg protein⁻¹ indicating a decrease of around threefold.

In conclusion, the associated *Pain-I* alleles *Pain_SA* and *Pain_DA*, which showed in 3D-analysis a charge effect of the sucrose binding site, which might lead to reduced conversion of sucrose, did not display strong physiological differences compared to the other alleles of the corresponding genotypes. The ideal biochemical characteristics outlined in Figure 4.2 for superior allelic invertases associated with superior chips quality were not clearly recognized for vacuolar invertase alleles.

The 30°C assay of *invGE* alleles (section 3.2.2.3, Table 3.2.13) showed that the two associated *invGE* alleles *E_SA* and *E_TA*, which are not amino acid identical but are

characterized by the associated SNP 1103*, displayed similar K_m values of approximately 20mM. From the tetraploid cultivar ‘Satina’ three alleles E_SA , E_SNI , and E_SN3 were analyzed. Regarding the enzyme affinity, E_SN3 displayed the lowest sucrose affinity with a K_m value of approximately 24mM, whilst the allele E_SNI showed the highest substrate affinity with a K_m value of around 17mM. The K_m value of the associated allele E_SA of 20mM was in between and was significantly different from E_SN3 .

From the tetraploid cultivar ‘Theresa’ two alleles E_TA and E_TNI were analyzed. Neither the K_m values nor the v_{max} values of these alleles differed significantly from each other.

Compared to the other *invGE* alleles, the associated alleles E_SA and E_TA did not show strong K_m and v_{max} differences. The lowest K_m value was approximately 17mM (E_SNI), the highest K_m value was 24mM in the case of the allele E_PI8NI .

It could be shown by 3D-analysis that the charge of the putative sucrose binding site of the associated *invGE* alleles did not display any differences like observed for associated *Pain-I* alleles. This is in agreement with determined K_m values, which did not show strong physiological differences. Biochemical parameters of *invGE* alleles did not indicate the presence of obvious superior invertase alleles according to the assumption concerning K_m and v_{max} characteristics mentioned above (Figure 4.2).

Biochemical analysis of *invGF* alleles showed that substrate affinities ranged from approximately 12mM in case of F_SN2 and 21mM for the allele F_P40NI (section 3.2.2.3, Table 3.2.15). The alleles F_SN4 and F_DNI , which clustered phenetically together (section 3.2.1.2.8, Figure 3.2.11) as observed for the associated *invGE* alleles E_SA , E_DA , and E_TA , displayed K_m values of 18mM and 13mM, respectively. Within the tested ‘Satina’ *invGF* alleles F_SN4 revealed the lowest sucrose affinity, which differs significantly from F_SNI and F_SN2 .

The ‘Diana’ allele F_DNI , which is not identical at amino acid level to F_SN4 but belongs to the same outlying group, displayed a K_m value of 13mM. These differences between F_SN4 and F_DNI were significant, whilst the differences within the tested ‘Diana’ alleles were similar and not significant.

The allele F_SN4 , which is considered as a representative of the associated SSCP fragment *invGf-4d*, showed among the tested ‘Satina’ alleles the lowest sucrose affinity. 3D structural analysis of this allele showed that a charge change of the putative sucrose binding site occurred due to the allelic amino acid composition. The electrostatic potential of the site switched from positive to negative, possibly leading to less sucrose conversion reflected by a high K_m value as detected for the allele F_SN4 . To what extent this allele might contribute to

potato chips quality remains unclear since *invGF* is expressed in flowers and leaves and to date no transcripts in tubers were detected.

Comparing the measured K_m values of vacuolar and cell wall-bound invertase isoforms revealed overall similar enzymatic characteristics. Nevertheless, there are K_m and v_{max} values of single *Pain-1*, *invGE*, or *invGF* alleles, which are distinct from the average alleles. Even though the biochemical analysis of *Pain-1*, *invGE*, or *invGF* alleles in the heterologous system yeast revealed only slight allelic differences regarding substrate affinity and substrate conversion, no statement can be made concerning the *in vivo* situation in potato.

The measured invertase affinities towards the substrate sucrose are in general agreement with previously detected K_m values. FRIDMAN ET AL. (2004) measured substrate affinities between approximately 5 and 12mM for a functional cell wall-bound tomato invertase isoform expressed in yeast. The *Arabidopsis* soluble invertase *INV2* displayed a K_m of 12mM (TANG ET AL., 1996). The substrate affinity for a soluble acid invertase from *Triticum aestivum* was set at 19.6mM (KRISHNAN ET AL., 1985).

Convincing evidence that the invertase assay is suitable for the detection of the corresponding enzyme activities is the fact that, concerning the yeast wild type invertase, a sucrose affinity of approximately 22mM was measured. GASCON ET AL. (1968) determined the K_m value of yeast invertase at 25mM, which is nearly identical to the K_m value detected in this study.

Documented K_m values for plant invertases as described above from the BRENDA database (<http://www.brenda-enzymes.org>) were similar to the measured values in this study.

4.4 Potato invertase isoforms and corresponding alleles display a large structural and functional variation and are interesting candidate genes in the trait potato chips quality

The identification of alleles of five different potato invertase genes revealed high allelic diversity within the selected six genotypes. However, there are certain alleles identified shared by more than one genotype, representing basic invertase alleles. It is remarkable that invertase alleles identical at amino acid level displayed synonymous polymorphisms in their nucleotide sequences. These ‘silent’ SNPs not leading to any amino acid exchanges might influence mRNA secondary structure, stability, translational efficiency, and protein folding leading to altered enzyme activity. Possible effects of synonymous SNPs were not subject of this study but the further investigation of these SNPs will clarify if there is a connection between the latter and the trait potato chips quality.

3D-modelling of the alleles gave first insights in possible structural and charge effects regarding amino acid differences between the alleles. It was found that alleles associated with superior potato chips quality showed charge changes in the putative sucrose binding site leading to a putative lower conversion of sucrose into the reducing sugars glucose and fructose. As these reducing sugars are known to interfere with potato chips quality, lower conversion would imply a better chips phenotype. To test, whether allele specific amino acid differences lead to altered enzyme properties, invertase assays were performed in the heterologous system yeast. Determining the enzyme’s affinity to sucrose (K_m) and the maximal rate of substrate conversion (v_{max}) showed that vacuolar and cell wall-bound invertase alleles displayed no differences that could explain the effect on chips quality. The observed differences might reflect the slight differences present *in vivo* since natural occurring alleles were analyzed and no mutant, wild accession, or other impaired alleles were tested like done for tomato (FRIDMAN ET AL., 2004). Genotype dependent alleles were detected, which displayed extreme values regarding their K_m and v_{max} values. The associated alleles were similar to not associated ones, indicating that K_m and v_{max} values do not seem to be relevant for possible reduced sucrose conversion into the sugars glucose and fructose and, therefore, better chips quality. Convincing evidence that the heterologous system yeast was suitable for assaying potato invertase activity was the analysis of two nucleotide variants of the *Pain-1* allele *Pain_SN* (section 3.1.2.3, Tables 3.1.20 and 3.1.21). The variability ranged from one to three nucleotide differences. It was shown that all nucleotide versions displayed similar K_m and v_{max} values, showing that a codon usage independent comparison of different alleles was possible.

Additionally, expression analysis revealed genotype dependent transcription patterns of invertase alleles. Associated alleles showed transcriptional changes for *Pain_SA* and *Pain_P18N1* compared to their genomic distribution. The latter is amino acid identical to *Pain_DA*, which showed no specific expression pattern. The transcript levels of associated *invGE* and the putative associated *invGF* alleles in leaves and flowers, respectively, were similar to their genomic dosages.

Invertase is functional as a complex of allelic subunits, thus it might be possible that allelic structural differences on the enzyme's surface, charge differences of the molecule as well as changed ratios of subunits due to transcriptional and translational regulation, lead to altered enzyme complexes less functional than in other genotypes with an inferior potato chips quality phenotype.

Another regulation of invertase activity is inhibition mediated by the endogenous invertase inhibitor (SCHWIMMER ET AL., 1961; PRESSEY, 1967; EWING AND MCADOO, 1971; WEIL ET AL., 1994; HEIBGES ET AL., 2003) or inhibition from a potato lectin (ISLA ET AL., 1991). The invertase inhibitor forms a non-dissociable complex with the enzyme, whilst the lectin is a dissociable inhibitor. Assuming allelic variants of invertase inhibitors acting differentially on allelic subunits of a putative invertase enzyme complex might lead to altered invertase activity. Also the potato lectin might act dependent on the allelic composition of an invertase complex in different ways to modulate invertase activity.

Association analysis of *Pain-I* allelic fragments revealed LD with allelic fragments of two other genes on chromosome III (LI ET AL., 2008). The first gene of this associated haplotype block located 6cM in the distal of *Pain-I* is the plastidic L-type α -Glucan phosphorylase (*Stp23*). The metabolic role of *Stp23* is the phosphoric degradation of starch. The second gene within the associated block is soluble starch synthase I (*SssI*) 14cM in the proximal of *Pain-I*. *SssI* plays a role in starch synthesis by connecting ADP-glucose to form linear amylose chains. It is possible that alleles of the other two associated genes might be responsible for or contribute to the studied trait potato chips quality. Also additive effects of alleles of all three loci *Pain-I*, *Stp23*, and *SssI* might be reasonable by their role in the carbohydrate metabolism leading to superior chips quality (OMHOLT ET AL., 2000; BIRCHLER ET AL., 2001).

In conclusion, the data presented in this study do not allow to a definite statement about the functional role of invertases on potato chips quality. However, it was shown that tremendous allelic variation of potato invertases exists, which leads to variation of transcript levels and biochemical parameters.

4.5 Future perspectives about the investigation of potato invertases and their natural variation

The results obtained in this thesis revealed important aspects about the diversity of invertase alleles and their regulation at transcriptional and biochemical level.

Considering the information available through this study, the question of ‘associated’ and ‘not associated’ alleles has to be addressed in a new manner. In the course of the work, no consistent functional characteristics of alleles statistically associated with better potato chips quality were determined. The nomenclature of association was based on the SSCP analysis by LI ET AL. (2005, 2008). Having now allele specific SNPs in hands, new alleles not detected so far by SSCP analysis but associated with superior chips quality can be analyzed in a larger population that allows screening for their impact on this trait. Being aware of allelic variability found in this study, new and up to now unidentified invertase alleles not detectable by SSCP analysis can be assessed by PCR based cloning. In consequence deeper biochemical and molecular characterization could enrich the understanding of allelic impact on potato chips quality.

Furthermore, allelic sequence variation strongly implies possible variations in regulatory domains like promoter sequences, enhancer-, and signalling domains. Additional studies of allelic promoter sequences and their influence on transcript levels might be useful to elucidate regulatory mechanisms of specific invertase alleles and identifying eQTLs as well as determining their impact on potato and the quantitative trait chips quality.

Another interesting aspect to follow is how invertase complexes might be build in potato, to what extend allelic monomers are involved in proper complex formation, how allelic monomer abundance due to transcriptional and translational regulations might influence enzyme complexes and, therefore, enzymatic activity.

The information gained by these additional approaches could help to draw a more detailed picture how invertases are regulated and to what extend allelic differences might contribute to a superior potato chips quality.

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Appendix

- A2** Map of the yeast expression vector 112 A1 NE.
- A3** Description of the amino acid exchanges using the alignment program Multalin version 5.4.1 (<http://bioinfo.genetoul.fr>).
- A3.1.1** Alignment of full-length amino acid sequences of *Pain-I* 'Satina' alleles.
- A3.1.2** Alignment of full-length amino acid sequences of *Pain-I* 'Diana' alleles.
- A3.1.3** Alignment of full-length amino acid sequences of *Pain-I* 'Theresa' alleles.
- A3.1.4** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_SA*.
- A3.1.5** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_SN*.
- A3.1.6** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_DA*.
- A3.1.7** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_DN1*.
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- A3.1.10** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_TN2*.
- A3.1.11** Alignment of full-length *Pain-I* nucleotide sequences obtained from all tetraploid genotypes.
- A3.1.12** Alignment of full-length amino acid sequences of *Pain-I* P18 alleles.
- A3.1.13** Alignment of full-length amino acid sequences of *Pain-I* P40 alleles.
- A3.1.14** Alignment of full-length amino acid sequences of *Pain-I* P54 alleles.
- A3.1.15** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-I* allele *Pain_P18N1*.
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- A3.1.18** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-1* allele *Pain_P40N2*.
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- A3.1.24** Alignment of the genomic *Pain-1* BAC insert sequence with the *Pain-1* cDNA sequence to identify the exon- and intron structure of the *Pain-1* gene.
- A3.1.25** Alignment of full-length nucleotide sequences from SNPs variants of the *Pain-1* allele *Pain_SN*.
- A3.2** Exception of alleles, from which only one full-length sequence was obtained.
- A3.2.1** Alignment of full-length amino acid sequences of *invGE* ‘Satina’ alleles.
- A3.2.2** Alignment of full-length amino acid sequences of *invGE* ‘Diana’ alleles.
- A3.2.3** Alignment of full-length amino acid sequences of *invGE* ‘Theresa’ alleles.
- A3.2.4** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_SA*.
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- A3.2.6** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_SN3*.
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- A3.2.8** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_DN1*.
- A3.2.9** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_TA*.

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- A3.2.10** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_TN1*.
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- A3.2.13** Alignment of full-length *invGE* nucleotide sequences obtained from all tetraploid genotypes.
- A3.2.14** Alignment of full-length amino acid sequences of *invGE* P18 alleles.
- A3.2.15** Alignment of full-length amino acid sequences of *invGE* P40 alleles.
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- A3.2.17** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_P18N1*.
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- A3.2.19** Alignment of full-length nucleotide sequences from SNPs variants of the *invGE* allele *E_P40N1*.
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- A3.2.24** Alignment of full-length *invGE* nucleotide sequences obtained from all tetraploid and diploid genotypes.
- A3.2.25** Alignment of full-length amino acid sequences of *invGF* 'Satina' alleles.
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- A3.2.28** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_SN1*.
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- A3.2.30** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_SN3*.
- A3.2.31** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_SN4*.
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- A3.2.33** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_TN1*.
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- A3.2.35** Alignment of full-length *invGF* nucleotide sequences obtained from all tetraploid genotypes.
- A3.2.36** Alignment of full-length amino acid sequences of *invGF* P18 alleles.
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- A3.2.39** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_P18N*.
- A3.2.40** Alignment of full-length nucleotide sequences from SNPs variants of the *invGF* allele *F_P40N2*.
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- A3.2.43** Alignment of full-length *invGF* nucleotide sequences obtained from all diploid genotypes.
- A3.2.44** Alignment of full-length *invGF* nucleotide sequences obtained from all tetraploid and diploid genotypes.
- A3.2.45** Alignment of full-length amino acid sequences of the *invGE* allele *E_SA* and *invGF* allele *F_SN4*.
- A3.3** Exception of alleles, from which only one full-length sequence was obtained.
- A3.3.1** Alignment of full-length amino acid sequences of *pCD111* ‘Satina’ alleles.
- A3.3.2** Alignment of full-length amino acid sequences of *pCD111* ‘Theresa’ alleles

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- A3.3.3** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD111* allele *CD111_S1*.
- A3.3.4** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD111* allele *CD111_S2*.
- A3.3.5** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD111* allele *CD111_T2*.
- A3.3.6** Alignment of full-length *pCD111* nucleotide sequences obtained from all tetraploid genotypes.
- A3.3.7** Alignment of full-length amino acid sequences of *pCD111* P54 alleles.
- A3.3.8** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD111* allele *CD111_P54_1*.
- A3.3.9** Alignment of full-length *pCD111* nucleotide sequences obtained from all diploid genotypes.
- A3.3.10** Alignment of full-length *pCD111* nucleotide sequences obtained from all tetraploid and diploid genotypes.
- A3.3.11** Alignment of full-length amino acid sequences of *pCD141* ‘Satina’ alleles.
- A3.3.12** Alignment of full-length amino acid sequences of *pCD141* ‘Diana’ alleles.
- A3.3.13** Alignment of full-length amino acid sequences of *pCD141* ‘Theresa’ alleles.
- A3.3.14** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_S2*.
- A3.3.15** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_S3*.
- A3.3.16** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_D1*.
- A3.3.17** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_T1*.
- A3.3.18** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_T2*.
- A3.3.19** Alignment of full-length *pCD141* nucleotide sequences obtained from all tetraploid genotypes.
- A3.3.20** Alignment of full-length amino acid sequences of *pCD141* P18 alleles.
- A3.3.21** Alignment of full-length amino acid sequences of *pCD141* P40 alleles.
- A3.3.22** Alignment of full-length amino acid sequences of *pCD141* P54 alleles.

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- A3.3.23** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_P18_1*.
- A3.3.24** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_P18_2*.
- A3.3.25** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_P40_1*.
- A3.3.26** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_P54_1*.
- A3.3.27** Alignment of full-length nucleotide sequences from SNPs variants of the *pCD141* allele *CD141_P54_2*.
- A3.3.28** Alignment of full-length *pCD141* nucleotide sequences obtained from all diploid genotypes.
- A3.3.29** Alignment of full-length *pCD141* nucleotide sequences obtained from all tetraploid and diploid genotypes.
- A3.3.30** Amino acid alignment of the *pCD111* BAC insert sequence and the P40 allele *CD111_P40_1*.
- A3.3.31** Amino acid alignment of the *pCD141* BAC insert sequence and the P40 allele *CD141_P40_1*.
- A3.3.32** Alignment of the genomic *pCD111* BAC insert sequence with the P40 *pCD111* cDNA sequence to identify the exon- and intron structure of the *pCD111* gene.
- A3.3.33** Alignment of the genomic *pCD141* BAC insert sequence with the P40 *pCD141* cDNA sequence to identify the exon- and intron structure of the *pCD141* gene.

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I am also grateful to Dr. Ute Achenbach who helped me a lot with reading my PhD work and besides being very critical regarding my English style, she also was able to encourage me. Her careful reading and useful comments improved this work noticeably.

Also thanks to the collaborators of this work: Dr. Alisdair Fernie, Dr. Adriano Nunes-Nesi and Pawel Durek (MPIMP/Golm). Dr. Alisdair Fernie and Dr. Adriano Nunes-Nesi gave me the possibility to perform the biochemical analysis in the laboratory in Golm. Pawel Durek modelled the invertase alleles and, therefore, special thanks to him. He was always open for my new ideas, for more and more allele models, and spared me with too many details about things I never heard anything before.

Thanks to Dr. Benjamin Stich (MPIZ/Köln) for calculation of the significance levels of the biochemical parameters.

Thanks to the ADIS service group, who really sequenced a lot of invertase alleles for me. Special thanks to Iris Bürst, who took so many times special care of my samples.

Thanks to my former colleagues Dr. Silke Schulze und Dr. Melanie Bartsch for so many nice lunch breaks, also for useful work discussions, scientific support and being there when things went wrong and I needed a good word to get balanced again.

Finally, I thank my family: my husband Michael, who was always there when melancholy thoughts and doubts were overwhelming leading me in way to believe in myself again. His endless patience was often working to full capacity but nevertheless always present. He also ensured that normal life went on, that our fridge was filled, the car was fuelled and that more dishes were inside the cupboard than outside. He also took affectionately care of our daughter and was able to dress up all the knights in the right way. Besides all this he also helped me bringing my work in a nice shape, formatting results of three years work. My daughter Berenike helped me with her naive point of view not understanding why things of ‘the adults’ are so important when the sun is reflecting itself in a water drop. My mother Petra was always there for me even if not physically present. We spent lots of hours at the phone convincing me that everything in life has its sense even if it sometimes takes a while until it becomes revealed. My mother in law Sabine supported us directly with her willingness taking care of our daughter when stress occupied me and my husband.

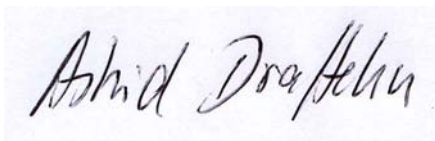
Die vorliegende Arbeit wurde am Max-Planck-Institut für Züchtungsforschung in Köln durchgeführt.

ERKLÄRUNG

Köln, 2009

Ich versichere, dass ich die von mir vorgelegte Dissertation selbständig angefertigt, die benutzten Quellen und Hilfsmittel vollständig angegeben und die Stellen der Arbeit – einschließlich Tabellen, Karten und Abbildungen – die anderen Werken im Wortlaut oder dem Sinn nach entnommen sind, in jedem Einzelfall als Entlehnung kenntlich gemacht habe; dass diese Dissertation noch keiner anderen Fakultät oder Universität zur Prüfung vorgelegen hat; dass sie noch nicht veröffentlicht worden ist sowie, dass ich eine solche Veröffentlichung vor Abschluss des Promotionsverfahrens nicht vornehmen werde.

Die Bestimmungen dieser Promotionsordnung sind mir bekannt. Die von mir vorgelegte Dissertation ist von PD Dr. Christiane Gebhardt betreut worden.

A handwritten signature in black ink on a light blue background. The signature is written in a cursive style and reads "Astrid Draffehn".

Astrid Draffehn

Lebenslauf

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Geburtsort Dessau
Staatsangehörigkeit deutsch
Familienstand verheiratet
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Kurse: Immunologie (Kurs für Master-Studiengang)
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Desert Ecology
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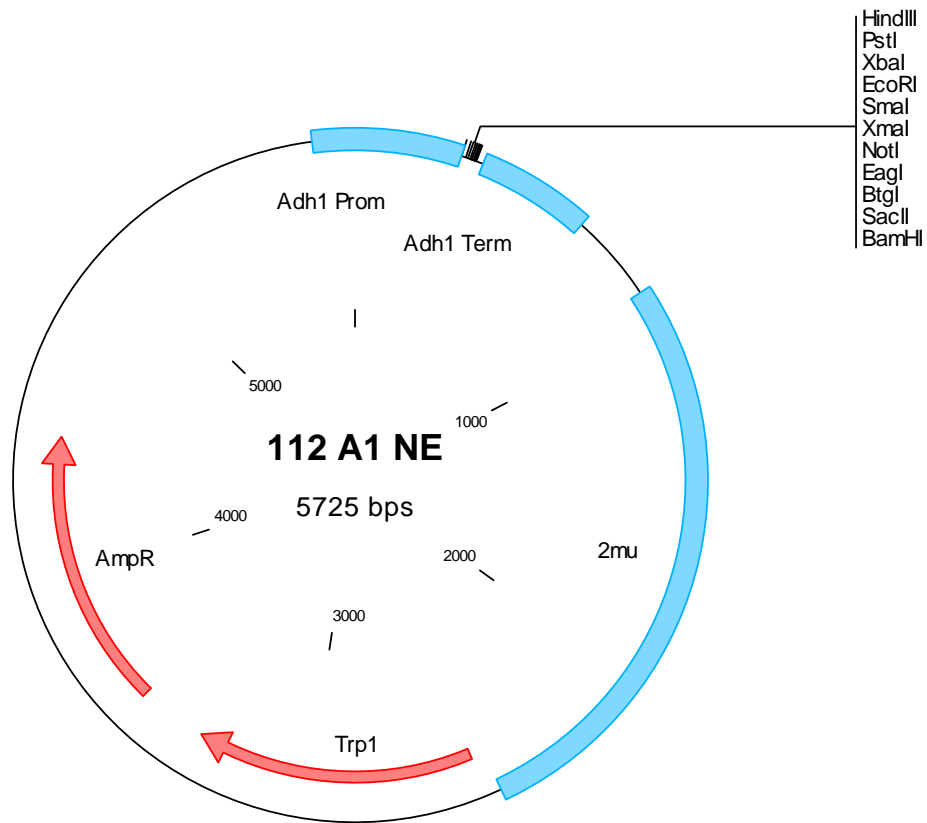
01/2001 – 08/2001 Praktisches Arbeiten in der Forschungsgruppe von Dr. Robert Webb,
University of Texas at El Paso, USA
(in Deutschland Anerkennung als Forschungsgruppenpraktikum)
Haupttätigkeit: Klonierung von bekannten und putativen
Metallothionein-Sequenzen aus Cyanobakterien in *E.coli*,
Überexpression der Metallothioneine, anschließende Protein-
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Parasitologie (Kurs für Doktoranden)
Desert Ecology
English
- 01/2001 – 08/2001 Praktisches Arbeiten in der Forschungsgruppe von Dr. Robert Webb, *University of Texas at El Paso*, USA
(in Deutschland Anerkennung als Forschungsgruppenpraktikum)
Haupttätigkeit: Klonierung von bekannten und putativen Metallothionein-Sequenzen aus Cyanobakterien in *E.coli*, Überexpression der Metallothioneine, anschließende Proteinaufreinigung sowie 3D-Protein-Struktur-Analyse
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- 03/2003 – 11/2003 Diplomarbeit, Thema: „Expression heterologer *psbA*-Gene in *Clamydomonas reinhardtii*“ an der *Martin-Luther-Universität Halle-Wittenberg* in der AG Prof. Dr. Udo Johanningmeier
- 11/2003 Verteidigung Diplomarbeit

Appendix A2

Map of the yeast expression vector 112 A1 NE.



Appendix A3

For aligning the amino acid and nucleotide sequences of the obtained alleles the following software was used:

Multalin version 5.4.1 (<http://bioinfo.genetoul.fr>)

Copyright I.N.R.A. France 1989, 1991, 1994, 1996

Multiple sequence alignment with hierarchical clustering

F. CORPET, 1988, Nucl. Acids Res., 16 (22), 10881-10890

Different matrixes were used for comparing the allelic sequences.

For all alignments except A3.1.24, A3.2.45, A3.3.32, and A3.3.33 the following parameters were used:

Symbol comparison table: **blosum62**
Gap weight: 12
Gap length weight: 2
Consensus levels: high=90% low=50%
Consensus symbols:
! is anyone of IV
\$ is anyone of LM
% is anyone of FY
is anyone of NDQEBZ

For aligning the sequences of A3.1.24, A3.2.45, A3.3.32, and A3.3.33 the following matrix was used:

Symbol comparison table: **dayhoff**
Gap weight: 5
Gap length weight: 0
Consensus levels: high=90% low=50%
Consensus symbols:
\$ is anyone of LM
% is anyone of FY
is anyone of BDENQZ

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SN1	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SN2	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SN3	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SN4	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SA	MATQYHSSYDP ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
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Pain_SA3	MATQYHSSYDP ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Pain_SA2	MATQYHSSYDP ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
Consensus	MATQYHSSYDL ENSASHYTFLPDQPD SGHRKSLKIISGIFLSSFLLL SVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWS#AMLSMQRTAYHFQPKNWHNDPNGPLYHKG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_SN1	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_SN2	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
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Pain_SN4	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_SA	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGIKDFRDPTTAWTGPQNGQWLL													
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Consensus	WYHLFYQYNPDSAIINGNITWGHAYSKDLIHWLYLPFAIVPDQWYDINGVNTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSOPLLLDWVKYKGNPVLVPPPGIGIKDFRDPTTAWTGPQNGQWLL													
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Pain_SN	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAYPGTGMWECVDFYPVST EKTNGLDTSYNGPGVKHVLKASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGYKYASKTFYDPKKQRRVWLG													
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	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
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Pain_SN1	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAELDIEASFVQKVALQGIIADHVGFSCSTSGGAASRGILGPFQVVIADQTLSELT													
Pain_SN2	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAELDIEASFVQKVALQGIIADHVGFSCSTSGGAASRGILGPFQVVIADQTLSELT													
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Pain_SA	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSYPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAYNGAARLFYFNNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_SA1	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSYPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAYNGAARLFYFNNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_SA3	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSYPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAYNGAARLFYFNNATGASVTASVKIWSLESANIRSFPLQDL													
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Consensus	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSYPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAYNGAARLFYFNNATGSSVTASVKIWSLESANIQSFPLQDL													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_TN1	MATQYHSSYDLENSASHYTFLPDQPDGHRKSLKIISGIFLSSFLLLSVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLNQRTAYHFQPQKNWMDPNGLYHKG													
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Pain_TN2_1	MATQYHSSYDLENSASHYTFLPDQPDGHRKSLKIISGIFLSSFLLLSVAFFPILNNQSPDLQSNRSPAPPSRGVSQGVSDKTFRDVYNASHVSYAWSNAMLNQRTAYHFQPQKNWMDPNGLYHKG													
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	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_TN1	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN1_1	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN1_2	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN1_3	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN2	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN2_1	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN2_2	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Pain_TN2_3	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
Consensus	WYHLFYQYNPDSAIWGNITWGHAYSKDLIHWLYLPFAHVPDQWYDINGVWTGSATILPDGQIHMLYTGDTDDYVQVQNLAYPTNLSPLLLDWVKYKGNPVLVPPPGIGYKDFRDPTTAWTGPQNGQWLL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_TN1	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN1_1	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN1_2	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN1_3	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN2	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN2_1	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN2_2	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_TN2_3	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Consensus	TIGSKIGKTGIALYYETSNTFSKLLDEVLAHVPGTGHWECVDFYPVSTEKTNGLDTSYNGPGVKHVLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_TN1	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN1_1	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN1_2	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN1_3	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN2	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN2_1	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN2_2	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Pain_TN2_3	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
Consensus	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQVNLQPGSIELLHVDSAAELDIASFEYDKVALQGIIEADHYGFSCSTSGGAASRGILGPFQVYVIADQTLSELT													
	521	530	540	550	560	570	580	590	600	610	620	630	639	
Pain_TN1	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGSSVTASYKINSLESANIQSFPLQDL													
Pain_TN1_1	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGSSVTASYKINSLESANIQSFPLQDL													
Pain_TN1_2	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGSSVTASYKINSLESANIQSFPLQDL													
Pain_TN1_3	PYYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGSSVTASYKINSLESANIQSFPLQDL													
Pain_TN2	PYYFFISKGADGRAETHFCADQTRSSEAPGVAKRVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGASVTASYKINSLESANIRSFPLQDL													
Pain_TN2_1	PYYFFISKGADGRAETHFCADQTRSSEAPGVAKRVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGASVTASYKINSLESANIRSFPLQDL													
Pain_TN2_2	PYYFFISKGADGRAETHFCADQTRSSEAPGVAKRVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGASVTASYKINSLESANIRSFPLQDL													
Pain_TN2_3	PYYFFISKGADGRAETHFCADQTRSSEAPGVAKRVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGASVTASYKINSLESANIRSFPLQDL													
Consensus	PYYF%ISKGADGRAETHFCADQTRSSEAPGVAKrVYGSSVPVLDGEKHSRLLVDHSIYESFAQGGRTVITSRIYPTKAVNGAARLFVFNATGASVTASYKINSLESANIRSFPLQDL													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_SA	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_SA1	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_SA3	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_SA2	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_SA	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCCAGTCACCGGACTTGACAGAGTAACTCCCGTTGCCGGGCCACCCTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGAGCTTTTCGAGA													
Pain_SA1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCCAGTCACCGGACTTGACAGAGTAACTCCCGTTGCCGGGCCACCCTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGAGCTTTTCGAGA													
Pain_SA3	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCCAGTCACCGGACTTGACAGAGTAACTCCCGTTGCCGGGCCACCCTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGAGCTTTTCGAGA													
Pain_SA2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCCAGTCACCGGACTTGACAGAGTAACTCCCGTTGCCGGGCCACCCTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGAGCTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCCAGTCACCGGACTTGACAGAGTAACTCCCGTTGCCGGGCCACCCTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGAGCTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_SA	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_SA1	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_SA3	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_SA2	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Consensus	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_SA	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_SA1	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_SA3	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_SA2	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_SA	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACACTTATCTGA													
Pain_SA1	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACACTTATCTGA													
Pain_SA3	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACACTTATCTGA													
Pain_SA2	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACACTTATCTGA													
Consensus	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_SA	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACTACTGCTTGAGCCGACCCCAAATGGGCAATGGCTTTTA													
Pain_SA1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACTACTGCTTGAGCCGACCCCAAATGGGCAATGGCTTTTA													
Pain_SA3	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACTACTGCTTGAGCCGACCCCAAATGGGCAATGGCTTTTA													
Pain_SA2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACTACTGCTTGAGCCGACCCCAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACTACTGCTTGAGCCGACCCCAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_SA	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCCAACCTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_SA1	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCCAACCTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_SA3	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCCAACCTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_SA2	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCCAACCTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCCAACCTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_SA	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_SA1	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_SA3	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_SA2	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_SA	GACAAGAACAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCCAAGAACAACGAAGAGTACTGTGGGGA													
Pain_SA1	GACAAGAACAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCCAAGAACAACGAAGAGTACTGTGGGGA													
Pain_SA3	GACAAGAACAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCCAAGAACAACGAAGAGTACTGTGGGGA													
Pain_SA2	GACAAGAACAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCCAAGAACAACGAAGAGTACTGTGGGGA													
Consensus	GACAAGAACAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCCAAGAACAACGAAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_SA	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Pain_SA1	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Pain_SA3	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Pain_SA2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Consensus	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_SA	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Pain_SA1	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Pain_SA3	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Pain_SA2	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Consensus	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_SA	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAAAAGCTATCTGAGCTAACG													
Pain_SA1	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAAAAGCTATCTGAGCTAACG													
Pain_SA3	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAAAAGCTATCTGAGCTAACG													
Pain_SA2	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAAAAGCTATCTGAGCTAACG													
Consensus	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAAAAGCTATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_SA	CCAGTTTACTTCTACATTTCTAAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACCAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_SA1	CCAGTTTACTTCTACATTTCTAAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACCAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_SA3	CCAGTTTACTTCTACATTTCTAAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACCAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_SA2	CCAGTTTACTTCTACATTTCTAAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACCAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Consensus	CCAGTTTACTTCTACATTTCTAAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACCAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_SA	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Pain_SA1	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Pain_SA3	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Pain_SA2	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Consensus	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_SA	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_SA1	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_SA3	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_SA2	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Consensus	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_SN	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
Pain_SN2	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
Pain_SN1	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
Pain_SN3	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
Pain_SN4	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
Consensus	ATG	GCC	ACG	CAG	TAC	CATT	CC	AGT	TAT	GAC	CT	TG	GAA	AACT
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_SN	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
Pain_SN2	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
Pain_SN1	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
Pain_SN3	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
Pain_SN4	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
Consensus	TCCTTTT	GCTTT	CTGTAG	CCTTCTTT	CCGATCCT	CA	CA	CA	CA	CA	CA	CA	CA	CA
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_SN	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
Pain_SN2	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
Pain_SN1	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
Pain_SN3	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
Pain_SN4	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
Consensus	TGTCGT	CAATGCTAG	TACAG	CTTTCTT	TATGCGTGGT	CCAATGCTATGCTTAGCT	GGCA	AA	GA	ACTGCTTACC	ATTT	CA	AC	CT
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_SN	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
Pain_SN2	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
Pain_SN1	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
Pain_SN3	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
Pain_SN4	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
Consensus	TGGTAT	CATCTTTTT	TATCA	ATA	CAATCCAG	ATTCAG	CTATT	TGGG	GA	AA	TAT	CAC	AT	GGGG
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_SN	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
Pain_SN2	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
Pain_SN1	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
Pain_SN3	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
Pain_SN4	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
Consensus	ACG	ATATA	AACGGT	GTCTG	GACTGGG	TCCG	TACC	ATCCT	AC	CCG	ATGGT	CAG	ATCAT	GATGCTTT
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_SN	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
Pain_SN2	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
Pain_SN1	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
Pain_SN3	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
Pain_SN4	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
Consensus	TCCTCT	CCCTCT	AGACTGGG	TCA	AGTACA	AGGCA	ACCCGG	TTCTG	GTTC	CT	CA	CC	CCG	CA
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_SN	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
Pain_SN2	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
Pain_SN1	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
Pain_SN3	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
Pain_SN4	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
Consensus	AC	ART	CGGG	CTA	AGATTGG	TAA	ACGGG	TATTG	CACTTGT	TTAT	GAA	ACTT	CA	RA
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_SN	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
Pain_SN2	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
Pain_SN1	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
Pain_SN3	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
Pain_SN4	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
Consensus	TT	AC	CCG	G	TAT	CA	CT	G	AAAA	AA	CA	AC	CGG	T
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_SN	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
Pain_SN2	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
Pain_SN1	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
Pain_SN3	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
Pain_SN4	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
Consensus	G	A	CA	AG	A	CA	AA	T	GG	A	CA	CC	G	A
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_SN	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
Pain_SN2	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
Pain_SN1	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
Pain_SN3	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
Pain_SN4	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
Consensus	TG	G	AT	TGGG	GA	AA	CT	G	AT	AG	TGA	AT	CT	G
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_SN	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
Pain_SN2	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
Pain_SN1	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
Pain_SN3	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
Pain_SN4	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
Consensus	TT	G	AA	AG	CT	T	AA	G	A	G	C	G	G	T
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_SN	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
Pain_SN2	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
Pain_SN1	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
Pain_SN3	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
Pain_SN4	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
Consensus	G	CT	CA	GG	GA	AT	TA	AG	C	AG	AT	CA	T	G
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_SN	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
Pain_SN2	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
Pain_SN1	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
Pain_SN3	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
Pain_SN4	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
Consensus	CC	AG	TT	T	ACT	TT	CT	AA	AG	GA	CT	GA	T	GG
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_SN	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
Pain_SN2	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
Pain_SN1	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
Pain_SN3	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
Pain_SN4	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
Consensus	AC	GG	T	GA	AA	AA	CA	TT	CG	AT	G	AG	AT	T
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_SN	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G
Pain_SN2	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G
Pain_SN1	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G
Pain_SN3	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G
Pain_SN4	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G
Consensus	C	G	T	T	T	CA	CA	AT	G	CC	AC	AG	G	G

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
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Pain_DA	ATGGCCACGCAGTACCATTCCAGTTATGACCCGGAAACTCCGCCTCCCATTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTT													
Pain_DA2	ATGGCCACGCAGTACCATTCCAGTTATGACCCGGAAACTCCGCCTCCCATTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTT													
Pain_DA1	ATGGCCACGCAGTACCATTCCAGTTATGACCCGGAAACTCCGCCTCCCATTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTT													
Consensus	ATGGCCACGCAGTACCATTCCAGTTATGACCCGGAAACTCCGCCTCCCATTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
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Pain_DA	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGAGTAATCCCCTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTTCGAGA													
Pain_DA2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGAGTAATCCCCTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTTCGAGA													
Pain_DA1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGAGTAATCCCCTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGAGTAATCCCCTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
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Pain_DA	TGTCGTCATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTGCCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_DA2	TGTCGTCATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTGCCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_DA1	TGTCGTCATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTGCCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Consensus	TGTCGTCATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTGCCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_DA	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DA2	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DA1	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCTTTTGCCATGGTTCCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCTTTTGCCATGGTTCCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
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Pain_DA	ACGATATTACGGTGTCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DA2	ACGATATTACGGTGTCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DA1	ACGATATTACGGTGTCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Consensus	ACGATATTACGGTGTCTGGACTGGGTCCGCCACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
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Pain_DA	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTATCAAGGACTTTAGAGACCCGACCCTGCTTGGACCGGACCCCAAATGGGCAATGGCTTTTA													
Pain_DA2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTATCAAGGACTTTAGAGACCCGACCCTGCTTGGACCGGACCCCAAATGGGCAATGGCTTTTA													
Pain_DA1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTATCAAGGACTTTAGAGACCCGACCCTGCTTGGACCGGACCCCAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTATCAAGGACTTTAGAGACCCGACCCTGCTTGGACCGGACCCCAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
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Pain_DA	ACAATCGGGTCTAAGATTGGTAACCGGGTATTGCGCTTGTATTGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_DA2	ACAATCGGGTCTAAGATTGGTAACCGGGTATTG Ca CTTGTATTGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_DA1	ACAATCGGGTCTAAGATTGGTAACCGGGTATTG Ca CTTGTATTGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTAT AG GGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAACCGGGTATTG Ca CTTGTATTGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATG AG GGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
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Pain_DA	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_DA2	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_DA1	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
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Pain_DA	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAAGAACAACGAAGAGTACTGTGGGGA													
Pain_DA2	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAAGAACAACGAAGAGTACTGTGGGGA													
Pain_DA1	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAAGAACAACGAAGAGTACTGTGGGGA													
Consensus	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAAGAACAACGAAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
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Pain_DA	TGGATTGGGGAACCTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Pain_DA2	TGGATTGGGGAACCTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Pain_DA1	TGGATTGGGGAACCTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Consensus	TGGATTGGGGAACCTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
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Pain_DA	TTGAAGCTTAAGAGTGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
Pain_DA2	TTGAAGCTTAAGAGTGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
Pain_DA1	TTGAAGCTTAAGAGTGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
Consensus	TTGAAGCTTAAGAGTGGGTGATCCTATTGTTAAGCAAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_DA	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTGGTGTCGTTGTAATTGCTGATCAAAGCTATCTGAGCTAACG													
Pain_DA2	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTGGTGTCGTTGTAATTGCTGATCAAAGCTATCTGAGCTAACG													
Pain_DA1	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTGGTGTCGTTGTAATTGCTGATCAAAGCTATCTGAGCTAACG													
Consensus	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTGGTGTCGTTGTAATTGCTGATCAAAGCTATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_DA	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCCGTGTTAG													
Pain_DA2	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCCGTGTTAG													
Pain_DA1	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCCGTGTTAG													
Consensus	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCCGTGTTAG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_DA	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Pain_DA2	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Pain_DA1	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
Consensus	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCCAAGGAGGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCACGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_DA	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCTTCCGTCAAGATTTGGTCACCTTGAGTCGGCTAATATTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_DA2	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCTTCCGTCAAGATTTGGTCACCTTGAGTCGGCTAATATTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_DA1	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCTTCCGTCAAGATTTGGTCACCTTGAGTCGGCTAATATTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Consensus	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCTTCCGTCAAGATTTGGTCACCTTGAGTCGGCTAATATTCGATCCTTCCCCTTGCAAGACTTGTAAC													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_DM1	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Pain_DM1_4	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Pain_DM1_3	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Pain_DM1_1	ATGGCCACGCGATACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Pain_DM1_2	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Pain_DM1_5	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAARAACTCCGCCTCCCATTTACACATTCTCTCCGGATCAACCCGATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_DM1	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_DM1_4	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_DM1_3	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_DM1_1	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_DM1_2	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_DM1_5	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Consensus	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_DM1	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_DM1_4	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_DM1_3	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_DM1_1	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_DM1_2	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_DM1_5	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Consensus	TGTGCTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAAATGCTATGCTTAGCTGGCAAGAGAACTGCTTACCATTTTCACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_DM1	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DM1_4	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DM1_3	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DM1_1	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DM1_2	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_DM1_5	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAAATCCAGATTACGCTATTTGGGGAARATACATATGGGGCCATGCCGATCCAAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_DM1	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DM1_4	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DM1_3	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DM1_1	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DM1_2	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_DM1_5	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Consensus	ACGATATAAACGGGTGCTCGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_DM1	TCCTCTCCTTTCTAGACTGGGTCAAGTACAAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCATGGCTTTTA													
Pain_DM1_4	TCCTCTCCTTTCTAGACTGGGTCAAGTACAAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCATGGCTTTTA													
Pain_DM1_3	TCCTCTCCTTTCTAGACTGGGTCAAGTACAAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTTGGTGTCAAG													

		1	10	20	30	40	50	60	70	80	90	100	110	120	130
	Pain_DM2	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_1	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_3	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_6	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_5	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_4	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Pain_DM2_2	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	Consensus	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
		131	140	150	160	170	180	190	200	210	220	230	240	250	260
	Pain_DM2	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_1	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_3	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_6	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_5	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_4	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Pain_DM2_2	T	C	C	T	T	T	G	C	T	T	T	C	T	T
	Consensus	T	C	C	T	T	T	G	C	T	T	T	C	T	T
		261	270	280	290	300	310	320	330	340	350	360	370	380	390
	Pain_DM2	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_1	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_3	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_6	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_5	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_4	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Pain_DM2_2	T	G	T	G	T	G	T	G	T	G	T	G	T	G
	Consensus	T	G	T	G	T	G	T	G	T	G	T	G	T	G
		391	400	410	420	430	440	450	460	470	480	490	500	510	520
	Pain_DM2	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_1	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_3	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_6	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_5	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_4	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Pain_DM2_2	T	G	G	T	A	T	A	T	A	T	A	T	A	T
	Consensus	T	G	G	T	A	T	A	T	A	T	A	T	A	T
		521	530	540	550	560	570	580	590	600	610	620	630	640	650
	Pain_DM2	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_1	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_3	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_6	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_5	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_4	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Pain_DM2_2	A	C	G	A	T	T	A	A	C	G	G	T	G	T
	Consensus	A	C	G	A	T	T	A	A	C	G	G	T	G	T
		651	660	670	680	690	700	710	720	730	740	750	760	770	780
	Pain_DM2	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_1	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_3	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_6	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_5	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_4	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Pain_DM2_2	T	C	C	T	C	T	T	A	G	A	C	T	T	A
	Consensus	T	C	C	T	C	T	T	A	G	A	C	T	T	A
		781	790	800	810	820	830	840	850	860	870	880	890	900	910
	Pain_DM2	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_1	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_3	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_6	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_5	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_4	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Pain_DM2_2	A	C	A	T	C	G	G	T	C	A	A	A	C	G
	Consensus	A	C	A	T	C	G	G	T	C	A	A	A	C	G
		911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
	Pain_DM2	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_1	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_3	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_6	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_5	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_4	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Pain_DM2_2	T	T	A	C	C	G	T	A	C	G	A	A	A	A
	Consensus	T	T	A	C	C	G	T	A	C	G	A	A	A	A
		1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
	Pain_DM2	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_1	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_3	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_6	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_5	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_4	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Pain_DM2_2	G	A	C	A	A	G	A	C	A	A	A	A	C	A
	Consensus	G	A	C	A	A	G	A	C	A	A	A	A	C	A
		1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
	Pain_DM2	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_1	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_3	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_6	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_5	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_4	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Pain_DM2_2	T	G	G	A	T	T	A	C	A	A	A	A	C	A
	Consensus	T	G	G	A	T	T	A	C	A	A	A	A	C	A
		1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
	Pain_DM2	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_1	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_3	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_6	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_5	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_4	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Pain_DM2_2	T	T	A	A	G	A	C	T	T	A	A	A	C	A
	Consensus	T	T	A	A	G	A	C	T	T	A	A	A	C	A
		1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
	Pain_DM2	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_1	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_3	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_6	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_5	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_4	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Pain_DM2_2	G	C	T	C	A	G	G	A	A	T	T	G	A	A
	Consensus	G	C	T	C	A	G	G	A	A	T	T	G	A	A
		1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
	Pain_DM2	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_1	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_3	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_6	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_5	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_4	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Pain_DM2_2	C	A	G	T	T	A	C	T	T	C	A	A	A	A
	Consensus	C	A	G	T	T	A	C	T	T	C	A	A	A	A
		1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
	Pain_DM2	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_1	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_3	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_6	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_5	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_4	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Pain_DM2_2	A	C	G	G	T	A	A	A	A	C	A	T	T	A
	Consensus	A	C	G	G	T	A	A	A	A	C	A	T	T	A
		1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
	Pain_DM2	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_1	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_3	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_6	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_5	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_4	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Pain_DM2_2	C	G	T	T	T	A	C	A	A	T	T	G	A	A
	Consensus														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACC T GATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_TN1_2	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACC T GATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_TN1_3	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACC C GATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACTCCGCTCCCATTTACACATTCCTCCCGGATCAACC b .GATTCCGGCCACCGGAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_TN1_2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_TN1_3	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_TN1_2	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_TN1_3	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Consensus	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_TN1_2	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_TN1_3	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Pain_TN1_2	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Pain_TN1_3	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Consensus	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGAGCCCCAAATGGGCAATGGCTTTTA													
Pain_TN1_2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGAGCCCCAAATGGGCAATGGCTTTTA													
Pain_TN1_3	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGAGCCCCAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGAGCCCCAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_TN1_2	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_TN1_3	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAGAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_TN1_2	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAGAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_TN1_3	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAGAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATACGGCCCGGGTGTAAGAGCATGTGTTAAAGCAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	GACAAGAA C AATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
Pain_TN1_2	GACAAGAA C AATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
Pain_TN1_3	GACAAGAA C AATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
Consensus	GACAAGAA b AATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCAGAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAGAA													
Pain_TN1_2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCAGAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAGAA													
Pain_TN1_3	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCAGAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAGAA													
Consensus	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCAGAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAGAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	TTGAAGCTTAAGAGCGGGTGATCCTATTGT T AAGCAAGTCAATCTTCACACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Pain_TN1_2	TTGAAGCTTAAGAGCGGGTGATCCTATTGT T AAGCAAGTCAATCTTCACACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Pain_TN1_3	TTGAAGCTTAAGAGCGGGTGATCCTATTGT T AAGCAAGTCAATCTTCACACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
Consensus	TTGAAGCTTAAGAGCGGGTGATCCTATTGT b AAGCAAGTCAATCTTCACACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAAGCCTCATTGAGTGGACAAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
Pain_TN1_2	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
Pain_TN1_3	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
Consensus	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	CCAGTTTACTTCTACATTTCTAAGGAGGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_TN1_2	CCAGTTTACTTCTACATTTCTAAGGAGGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_TN1_3	CCAGTTTACTTCTACATTTCTAAGGAGGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Consensus	CCAGTTTACTTCTACATTTCTAAGGAGGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAAGATCCTCAGAGGCTCCGGGAGTTGCTAAACAAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	ACGGTGAAAAACATT C GATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGGAGAGACAGTCATACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAG C GCACGACTCTT													
Pain_TN1_2	ACGGTGAAAAACATT C GATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGGAGAGACAGTCATACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAG C GCACGACTCTT													
Pain_TN1_3	ACGGTGAAAAACATT C GATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGGAGAGACAGTCATACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAG C GCACGACTCTT													
Consensus	ACGGTGAAAAACATT b GATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGGAGAGACAGTCATACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAG b GCACGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_TN1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN1_1	CGTTTTCAACAAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCTTGCAGAGCTTGTA													
Pain_TN1_2	CGTTTTCAACAAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCTTGCAGAGCTTGTA													
Pain_TN1_3	CGTTTTCAACAAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCTTGCAGAGCTTGTA													
Consensus	CGTTTTCAACAAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCTTGCAGAGCTTGTA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCCTCCATTACACATTCTCCCGGATCAACCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_TN2_1	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCCTCCATTACACATTCTCCCGGATCAACCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_TN2_2	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCCTCCATTACACATTCTCCCGGATCAACCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Pain_TN2_3	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCCTCCATTACACATTCTCCCGGATCAACCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAAACTCCGCCTCCATTACACATTCTCCCGGATCAACCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGACAGAGTAATCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_TN2_1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGACAGAGTAATCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_TN2_2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAATTCACCGGACTTGACAGAGTAATCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_TN2_3	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGACAGAGTAATCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGACAGAGTAATCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_TN2_1	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_TN2_2	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_TN2_3	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Consensus	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	TGGTATCATCTTTTTATCAATACATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGGCATGGTTCCTGATCAATGGT													
Pain_TN2_1	TGGTATCATCTTTTTATCAATACATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGGCATGGTTCCTGATCAATGGT													
Pain_TN2_2	TGGTATCATCTTTTTATCAATACATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGGCATGGTTCCTGATCAATGGT													
Pain_TN2_3	TGGTATCATCTTTTTATCAATACATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGGCATGGTTCCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGGCATGGTTCCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGGGTACCCACCACTTATCTGA													
Pain_TN2_1	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGGGTACCCACCACTTATCTGA													
Pain_TN2_2	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGGGTACCCACCACTTATCTGA													
Pain_TN2_3	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGGGTACCCACCACTTATCTGA													
Consensus	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGGGTACCCACCACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCCTCCACCAGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGGACCCCAAATGGGCAATGGCTCTTA													
Pain_TN2_1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCCTCCACCAGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGGACCCCAAATGGGCAATGGCTCTTA													
Pain_TN2_2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCCTCCACCAGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGGACCCCAAATGGGCAATGGCTCTTA													
Pain_TN2_3	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCCTCCACCAGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGGACCCCAAATGGGCAATGGCTCTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCCTCCACCAGGCATTGGTGTCAAGGACTTTAGAGACCCGACCCTGCTTGAGCCGGACCCCAAATGGGCAATGGCTCTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_TN2	ACATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTTGTTTATGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGAGTGTGTGGACT													
Pain_TN2_1	ACATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTTGTTTATGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGAGTGTGTGGACT													
Pain_TN2_2	ACATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTTGTTTATGAACCTTCAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGAGTGTGTGGACT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_DA	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_SA	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_DM2	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_TN2	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_SN	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_TN1	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Pain_DM1	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
Consensus	AT	G	G	C	C	A	G	A	T	A	C	C	A	T
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_DA	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_SA	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_DM2	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_TN2	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_SN	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_TN1	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Pain_DM1	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
Consensus	TC	C	T	T	T	G	C	T	T	C	T	T	T	C
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_DA	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_SA	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_DM2	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_TN2	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_SN	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_TN1	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Pain_DM1	T	G	T	C	G	T	A	G	T	A	G	T	A	G
Consensus	T	G	T	C	G	T	A	G	T	A	G	T	A	G
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_DA	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_SA	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_DM2	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_TN2	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_SN	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_TN1	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Pain_DM1	T	G	T	A	T	A	T	A	T	A	T	A	T	A
Consensus	T	G	T	A	T	A	T	A	T	A	T	A	T	A
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_DA	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_SA	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_DM2	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_TN2	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_SN	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_TN1	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Pain_DM1	A	C	G	A	T	A	A	C	G	G	T	G	T	A
Consensus	A	C	G	A	T	A	A	C	G	G	T	G	T	A
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_DA	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_SA	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_DM2	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_TN2	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_SN	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_TN1	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Pain_DM1	T	C	T	C	T	C	T	A	G	A	C	T	A	G
Consensus	T	C	T	C	T	C	T	A	G	A	C	T	A	G
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_DA	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_SA	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_DM2	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_TN2	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_SN	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_TN1	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Pain_DM1	A	C	A	A	T	C	G	G	T	A	A	A	C	G
Consensus	A	C	A	A	T	C	G	G	T	A	A	A	C	G
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_DA	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_SA	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_DM2	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_TN2	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_SN	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_TN1	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Pain_DM1	T	T	A	C	C	G	G	T	A	C	G	A	C	A
Consensus	T	T	A	C	C	G	G	T	A	C	G	A	C	A
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_DA	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_SA	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_DM2	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_TN2	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_SN	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_TN1	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Pain_DM1	G	A	C	A	A	G	A	C	A	A	T	A	A	C
Consensus	G	A	C	A	A	G	A	C	A	A	T	A	A	C
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_DA	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_SA	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_DM2	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_TN2	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_SN	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_TN1	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_DM1	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Consensus	T	G	G	A	T	T	G	A	T	T	G	A	T	T
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_DA	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_SA	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_DM2	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_TN2	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_SN	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_TN1	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Pain_DM1	T	T	A	A	G	A	C	T	T	A	A	G	A	C
Consensus	T	T	A	A	G	A	C	T	T	A	A	G	A	C
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_DA	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_SA	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_DM2	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_TN2	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_SN	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_TN1	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Pain_DM1	G	C	T	C	A	G	G	A	A	T	T	G	A	G
Consensus	G	C	T	C	A	G	G	A	A	T	T	G	A	G
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_DA	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_SA	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_DM2	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_TN2	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_SN	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_TN1	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Pain_DM1	C	A	G	T	T	A	C	T	T	C	A	A	G	G
Consensus	C	A	G	T	T	A	C	T	T	C	A	A	G	G
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_DA	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_SA	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_DM2	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_TN2	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_SN	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_TN1	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_DM1	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Consensus	A	C	G	G	T	A	A	A	A	C	A	T	T	C
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_DA	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_SA	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_DM2	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_TN2	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_SN	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_TN1	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Pain_DM1	C	G	T	T	T	C	A	A	T	T	G	A	A	C
Consensus	C	G	T	T	T	C	A	A	T	T	G	A	A	C

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_P40N1	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N1_1	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N1_2	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N1_4	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N1_3	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N2	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNARSPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N2_1	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNARSPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Pain_P40N2_2	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNARSPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
Consensus	MATQYHSSYDPENSASHYTFLPDQHDSGHRKSLKIISGIFLSSLLLLSLVFFPILNNQSPDLQSNASHPAPPSRGVSQGVSDKTFRDVYNASHVSYAMSNAHLSMQRTAYHFQPQKNWMNDPNGPLYHKG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_P40N1	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N1_1	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N1_2	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N1_4	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N1_3	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N2	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N2_1	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P40N2_2	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
Consensus	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMVPDQWYDINGVMTGSATILPDGQIMHLYTGDTDDYVQVQNLAYPANLSDPLLLDWVKYKGNPVLVPPPGIGVKDFRDPTTAWTGPQNGQWLL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_P40N1	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N1_1	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N1_2	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N1_4	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N1_3	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N2	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N2_1	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Pain_P40N2_2	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
Consensus	TIGSKIGKTGIALVYETSNTFSFKLLDEVLHAYPGTGMECVDFYPVSTKTNGLDTSYNGPGVKHYLKAASLDNKKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKLDYGKYYASKTFYDPKKQRRVLWG													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_P40N1	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N1_1	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N1_2	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N1_4	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N1_3	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N2	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N2_1	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Pain_P40N2_2	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
Consensus	WIGETDSESADLQKGWASVQSIPTVLYDKKTRTHLLQWPVEEIESLRAGDPIVKQANLQPGSIELLHVDSAAELDIASFEYDKVALQGIEADHYGFSCSTSGGAASRGILGPFQVYVVIADQTLSELT													
	521	530	540	550	560	570	580	590	600	610	620	630	639	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_P40N1	PYYFYISKGADGRAQTHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N1_1	PYYFYISKGADGRAQTHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N1_2	PYYFYISKGADGRAQTHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N1_4	PYYFYISKGADGRAQTHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N1_3	PYYFYISKGADGRAQTHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N2	PYYFYISKGADGRAETHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N2_1	PYYFYISKGADGRAETHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Pain_P40N2_2	PYYFYISKGADGRAETHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													
Consensus	PYYFYISKGADGRAETHFCADQTRSSVAPGVAKQVYGSSVPVLDGEKHSRLLVDHSIVESFAQGGRTVITSRIYPTKAYNGAARLFVFNATGASVTASVKIWSLESANIRSFPLQDL													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P54N	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Pain_P54N_1	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Pain_P54N_2	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Pain_P54N_4	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Pain_P54N_5	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Pain_P54N_3	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
Consensus	MATQYHSSYDLENSASHYTFLPDQPDSGHRKSLKIISGIFLSSFLLLVAFFPILNNQSPDLQSNRSPAPPSRGVSGQVSDKTFRDVYNASHVSYAMSNAMLSMQRTAYHFQPQKNWMDPNGLYHKG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P54N	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P54N_1	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P54N_2	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P54N_4	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P54N_5	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Pain_P54N_3	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
Consensus	WYHLFYQYNPDSAINGNITWGHAYSKDLIHWLYLPFAMYPDQMYDINGVMTGSATILPDGQIMHLYTGD TDDYVQVQNLAYPTNLSDP LLLDWVKYKGNPVLYPPP GIGVKDFRDPTTAWTGPQNGQWLL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P54N	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_P54N_1	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_P54N_2	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_P54N_4	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_P54N_5	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Pain_P54N_3	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
Consensus	TIGSKIGKTGIALVYETSNFTSFKLLDEV LHAVPGTGMWECYDFYPVSTEKTNGLDTSYNGPGVKHYLKASLDDNKQDHYAIGTYDLTKNKWTPDNPELDCGIGLKL DYGKYYASKTFYDPKKQRRVLMG													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P54N	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGTASRGILGPF GVVVIADQTLSELT													
Pain_P54N_1	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGTASRGILGPF GVVVIADQTLSELT													
Pain_P54N_2	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGASRGILGPF GVVVIADQTLSELT													
Pain_P54N_4	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGASRGILGPF GVVVIADQTLSELT													
Pain_P54N_5	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGASRGILGPF GVVVIADQTLSELT													
Pain_P54N_3	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGASRGILGPF GVVVIADQTLSELT													
Consensus	WIGETDSESADLQKGWASVQSIPTVLYDKKTGTHLLQWPVEEIESLRAGDPIVKQYNLQPGSIELLHYDSAAELDIASFEVDKVALQGIIEADHYGFSCSTSGGaASRGILGPF GVVVIADQTLSELT													
	521	530	540	550	560	570	580	590	600	610	620	630	639	
Pain_P54N	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGS SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Pain_P54N_1	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGS SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Pain_P54N_2	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGS SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Pain_P54N_4	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVYGS SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Pain_P54N_5	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVCG SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Pain_P54N_3	PVYFYISKGADGRAETHFCADQTRSSEAPGA AKQVYSS SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													
Consensus	PVYFYISKGADGRAETHFCADQTRSSEAPGVAKQVyg SVPVLDGEKHSRLLVDHSIVESFAQGGRVITSRIYPTKAVNGAARLFVFN NATGSSYTASVKINSLESANIQSFLQDL													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P18N1	-----													
Pain_P18N1_2	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
Pain_P18N1_1	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P18N1	-----													
Pain_P18N1_2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGCAGAGTAAGTCCCCTTCGCCGGCGCCGCCGTCAGAGGGTGTTCCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
Pain_P18N1_1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGCAGAGTAAGTCCCCTTCGCCGGCGCCGCCGTCAGAGGGTGTTCCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACACCAAGTCACCGGACTTGCAGAGTAAGTCCCCTTCGCCGGCGCCGCCGTCAGAGGGTGTTCCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P18N1	-----													
Pain_P18N1_2	TGTCGTCAATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
Pain_P18N1_1	TGTCGTCAATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
Consensus	TGTCGTCAATGCTAGTCACATTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P18N1	-----													
Pain_P18N1_2	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCCTTTGCCATGGTTCTGATCAATGGT													
Pain_P18N1_1	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCCTTTGCCATGGTTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCCTTTGCCATGGTTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_P18N1	-----													
Pain_P18N1_2	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCCTACCCACCAACTTATCTGA													
Pain_P18N1_1	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCCTACCCACCAACTTATCTGA													
Consensus	ACGATATTACGGGTGCTGGACTGGGTCCGCCACCATCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCCTACCCACCAACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_P18N1	-----													
Pain_P18N1_2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
Pain_P18N1_1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTATCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_P18N1	-----													
Pain_P18N1_2	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCAACCTTCAACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P18N1_1	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCAACCTTCAACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTTCAACCTTCAACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_P18N1	-----													
Pain_P18N1_2	TTTACCCGGTATCGACTGA AAAAACAAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCAAGTGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P18N1_1	TTTACCCGGTATCGACTGA AAAAACAAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCAAGTGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGA AAAAACAAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCAAGTGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_P18N1	-----													
Pain_P18N1_2	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAGAAACAAACGAGAGTACTGTGGGGA													
Pain_P18N1_1	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAGAAACAAACGAGAGTACTGTGGGGA													
Consensus	GACAAGAACAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGCATTTTATGACCCGAGAAACAAACGAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_P18N1	-----													
Pain_P18N1_2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCCAAGGACAGTGCTTTACGACAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Pain_P18N1_1	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTCCAAGGACAGTGCTTTACGACAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA</													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130

Pain_P18N2	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACCTCCGCTCCCATTAACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Pain_P18N2_1	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACCTCCGCTCCCATTAACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Pain_P18N2_3	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACCTCCGCTCCCATTAACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Pain_P18N2_2	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACCTCCGCTCCCATTAACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Consensus	ATGGCCACGCGAGTACCATTCCAGTTATGACCTGGAAAACCTCCGCTCCCATTAACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260

Pain_P18N2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_P18N2_1	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_P18N2_3	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_P18N2_2	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGACAGAGTAACTCCCGTTCGCCGGCGCCGCCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390

Pain_P18N2	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_P18N2_1	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_P18N2_3	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Pain_P18N2_2	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
Consensus	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTTCAACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520

Pain_P18N2	TGGTATCATCTTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_P18N2_1	TGGTATCATCTTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_P18N2_3	TGGTATCATCTTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_P18N2_2	TGGTATCATCTTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTTATCAATACAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650

Pain_P18N2	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Pain_P18N2_1	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Pain_P18N2_3	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Pain_P18N2_2	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
Consensus	ACGATATAAACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAAGTGCAAAATCTTGCGTACCCACCAACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780

Pain_P18N2	TCCTCTCCTTCTAGACTGGGTCA g GTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCA a AGACTTTAGAGACCCGACCACTGCTTGAGCCGGACCCCAAATGGGCAATGGCTTTTA													
Pain_P18N2_1	TCCTCTCCTTCTAGACTGGGTCA g GTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCA a AGACTTTAGAGACCCGACCACTGCTTGAGCCGGACCCCAAATGGGCAATGGCTTTTA													
Pain_P18N2_3	TCCTCTCCTTCTAGACTGGGTCA g GTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCA a AGACTTTAGAGACCCGACCACTGCTTGAGCCGGACCCCAAATGGGCAATGGCTTTTA													
Pain_P18N2_2	TCCTCTCCTTCTAGACTGGGTCA g GTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCA a AGACTTTAGAGACCCGACCACTGCTTGAGCCGGACCCCAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCA g GTACAAGGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCA a AGACTTTAGAGACCCGACCACTGCTTGAGCCGGACCCCAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910

Pain_P18N2	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACCTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P18N2_1	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACCTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P18N2_3	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACCTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P18N2_2	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACCTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAARACGGGTATTGCACCTTGTTTATGAARCTTCCAACCTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040

Pain_P18N2	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACG T ATGACTT													
Pain_P18N2_1	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACG T ATGACTT													
Pain_P18N2_3	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACG C ATGACTT													
Pain_P18N2_2	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACG T ATGACTT													
Consensus	TTTACCCGGTATCGACTGAAAAACAACCGGGTTGGACACATCATATAACGGCCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACG l ATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170

Pain_P18N2	GACAAGAATAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAGAACACGAGAGTACTGTGGGGA													
Pain_P18N2_1	GACAAGAATAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAGAACACGAGAGTACTGTGGGGA													
Pain_P18N2_3	GACAAGAATAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAGAACACGAGAGTACTGTGGGGA													
Pain_P18N2_2	GACAAGAATAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAGAACACGAGAGTACTGTGGGGA													
Consensus	GACAAGAATAAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAGAACACGAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300

Pain_P18N2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P18N2_1	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P18N2_3	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P18N2_2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Consensus	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430

Pain_P18N2	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCCG													
Pain_P18N2_1	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCCG													
Pain_P18N2_3	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCCG													
Pain_P18N2_2	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCCG													
Consensus	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGTCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCCG													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560

Pain_P18N2	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAACGCTATCTGAGCTAAGC													
Pain_P18N2_1	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAACGCTATCTGAGCTAAGC													
Pain_P18N2_3	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAACGCTATCTGAGCTAAGC													
Pain_P18N2_2	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAACGCTATCTGAGCTAAGC													
Consensus	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCGTTGTAATTGCTGATCAACGCTATCTGAGCTAAGC													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690

Pain_P18N2	CCAGTTTACT T CTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAAACAGATCCTCAGAGGCTCCGGGAGTTGCTAARACAGTTTATGGTAGTTCAGTACCCGTTGTG													
Pain_P18N2_1	CCAGTTTACT T CTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAAACAGATCCTCAGAGGCTCCGGGAGTTGCTAARACAGTTTATGGTAGTTCAGTACCCGTTGTG													
Pain_P18N2_3	CCAGTTTACT T CTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAAACAGATCCTCAGAGGCTCCGGGAGTTGCTAARACAGTTTATGGTAGTTCAGTACCCGTTGTG													
Pain_P18N2_2	CCAGTTTACT T CTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAAACAGATCCTCAGAGGCTCCGGGAGTTGCTAARACAGTTTATGGTAGTTCAGTACCCGTTGTG													
Consensus	CCAGTTTACT T CTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAAACAGATCCTCAGAGGCTCCGGGAGTTGCTAARACAGTTTATGGTAGTTCAGTACCCGTTGTG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820

Pain_P18N2	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCCTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGACAGTCAATACATCGCAATTTACCCACAAGGACAGTGAATGGAGCAGCAGACTCTT													
Pain_P18N2_1	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCCTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGACAGTCAATACATCGCAATTTACCCACAAGGACAGTGAATGGAGCAGCAGACTCTT													
Pain_P18N2_3	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCCTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGACAGTCAATACATCGCAATTTACCCACAAGGACAGTGAATGGAGCAGCAGACTCTT													
Pain_P18N2_2	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCCTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGACAGTCAATACATCGCAATTTACCCACAAGGACAGTGAATGGAGCAGCAGACTCTT													
Consensus	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCCTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGACAGTCAATACATCGCAATTTACCCACAAGGACAGTGAATGGAGCAGCAGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			

Pain_P18N2	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_P18N2_1	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_P18N2_3	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_P18N2_2	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAAC													
Consensus	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAAC													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P40N1	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
Pain_P40N1_2	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
Pain_P40N1_4	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
Pain_P40N1_3	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
Pain_P40N1_1	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
Consensus	ATGGCCACGCGAGTACCACTCAGGTTATGACCCGGAAAACTCCGCCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCCTCTC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P40N1	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P40N1_2	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P40N1_4	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P40N1_3	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P40N1_1	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCACCACCAAGTCACCGGACTTGC AAGTAAACGCCATTTCGCCGGCGCCGCGTC AAGAGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P40N1	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_P40N1_2	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_P40N1_4	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_P40N1_3	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Pain_P40N1_1	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
Consensus	TGTCGTCAATGCTAGTCA C GTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P40N1	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_P40N1_2	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_P40N1_4	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_P40N1_3	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Pain_P40N1_1	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAATCCAGATTCAAGTATTTGGGGAATATCACATGGGGCCATGCCGATCCAAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_P40N1	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
Pain_P40N1_2	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
Pain_P40N1_4	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
Pain_P40N1_3	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
Pain_P40N1_1	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
Consensus	ACGATATTACGGGTGCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAAGTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_P40N1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P40N1_2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P40N1_4	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P40N1_3	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P40N1_1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAAGCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_P40N1	AC AATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P40N1_2	AC AATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P40N1_4	AC GATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P40N1_3	AC AATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P40N1_1	AC AATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	AC aATCGGGTCTAAGATTGGTAAACCGGGTATTGCACCTTGTTTATGAACCTTCCAACTTCACAGCTTTAAACTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_P40N1	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P40N1_2	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P40N1_4	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P40N1_3	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P40N1_1	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGA AAAAACAAACAGCGGTTGGACACATCATATAACGGCCGGGTGTAAGCATGTGTTAAAGCAAGTTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_P40N1	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
Pain_P40N1_2	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
Pain_P40N1_4	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
Pain_P40N1_3	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
Pain_P40N1_1	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
Consensus	GACAAAGAACAAATGGACACCCGATACCCGGAATTGGATTGTGGAAATGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGAGACATTTTATGACCCGAAGAAACAAACGAGAGTGCTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_P40N1	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P40N1_2	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P40N1_4	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P40N1_3	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Pain_P40N1_1	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
Consensus	TGGATTGGGGAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTC AAGGACAGTGCTTTACGACAGAGAGACAGGACACATCTACTTCAGTGGCCAGTTGAGAAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_P40N1	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
Pain_P40N1_2	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
Pain_P40N1_4	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
Pain_P40N1_3	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
Pain_P40N1_1	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
Consensus	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAGGCCAATCTTCAACCAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_P40N1	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
Pain_P40N1_2	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
Pain_P40N1_4	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
Pain_P40N1_3	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
Pain_P40N1_1	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
Consensus	GCTCCAGGGAATTAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_P40N1	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
Pain_P40N1_2	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
Pain_P40N1_4	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
Pain_P40N1_3	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
Pain_P40N1_1	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
Consensus	CCAGTTTACTTCTACATTTCTA AAGGAGCTGATGGTCGAGCTCAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTA AACAAGTTTATGGTAGTTCAGTACCCG T TTGG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_P40N1	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
Pain_P40N1_2	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
Pain_P40N1_4	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
Pain_P40N1_3	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
Pain_P40N1_1	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
Consensus	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACCTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAGAACAGTCATAACATCACGAATTTACCCAAACAAGGCGAGTGATGGAGCAGCACGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_P40N1	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													
Pain_P40N1_2	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													
Pain_P40N1_4	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													
Pain_P40N1_3	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													
Pain_P40N1_1	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													
Consensus	CGTTTTCAACAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGATTTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCTTGC AAGACTTGTAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P40N2	-----													
Pain_P40N2_1	ATGGCCACGCGAGTACCCTCAAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTC													
Pain_P40N2_2	ATGGCCACGCGAGTACCCTCAAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTC													
Consensus	ATGGCCACGCGAGTACCCTCAAGTTATGACCCGGAAACTCCGCTCCCATTACACATTCTCCCGGATCAACACGATTCCGGCCACCGGAATCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P40N2	-----													
Pain_P40N2_1	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGTAACGCCCGTTCCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
Pain_P40N2_2	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGTAACGCCCGTTCCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCCTCAACAACCAAGTCACCGGACTTGCAAGTAACGCCCGTTCCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGTAAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P40N2	-----													
Pain_P40N2_1	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAACGGTCCATTGTACCACAGGGA													
Pain_P40N2_2	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAACGGTCCATTGTACCACAGGGA													
Consensus	TGTCGTCAATGCTAGTCACGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAACGGTCCATTGTACCACAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P40N2	-----													
Pain_P40N2_1	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Pain_P40N2_2	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATACAATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_P40N2	-----													
Pain_P40N2_1	ACGATATTACGGGGTCTGGACTGGGTCCGCTACCATCCTACCGGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAACTTATCTGA													
Pain_P40N2_2	ACGATATTACGGGGTCTGGACTGGGTCCGCTACCATCCTACCGGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAACTTATCTGA													
Consensus	ACGATATTACGGGGTCTGGACTGGGTCCGCTACCATCCTACCGGATGGTCAGATCATGATGCTTTATACTGGTGACACTGATGATTATGTGCAGTGCAAAATCTTGCGTACCCGCCAACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_P40N2	-----													
Pain_P40N2_1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCAAATGGGCAATGGCTTTTA													
Pain_P40N2_2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAGGCCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_P40N2	-----													
Pain_P40N2_1	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTCAAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_P40N2_2	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTCAAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACCTGTTTATGAACCTCAAACTTCACAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_P40N2	-----													
Pain_P40N2_1	TTTACCCGGTATCGACCGAAAAACAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_P40N2_2	TTTACCCGGTATCGACCGAAAAACAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACCGAAAAACAACGGGTTGGACACATCATATAACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACATAAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_P40N2	-----													
Pain_P40N2_1	GACAAGAACCAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACAACGAGAGTACTGTGGGGA													
Pain_P40N2_2	GACAAGAACCAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACAACGAGAGTACTGTGGGGA													
Consensus	GACAAGAACCAATGGACACCCGATAACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACAACGAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_P40N2	-----													
Pain_P40N2_1	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAAGGGATGGGCATCTGTACAGAGTATTCCAAGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Pain_P40N2_2	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAAGGGATGGGCATCTGTACAGAGTATTCCAAGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
Consensus	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAAGGGATGGGCATCTGTACAGAGTATTCCAAGGACAGTGCTTTACGACAGAAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_P40N2	-----													
Pain_P40N2_1	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGCCAACTTCAACCAGGTTGATTGAGCTACTCCATGTTGACTCAGC T GCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
Pain_P40N2_2	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGCCAACTTCAACCAGGTTGATTGAGCTACTCCATGTTGACTCAGC T GCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
Consensus	TTGAAGCTTAAGAGCGGGTGATCCTATTGTTAAGCAAGCCAACTTCAACCAGGTTGATTGAGCTACTCCATGTTGACTCAGC T GCAGAGTTGGATATAGAAGCCTCATTTGAAGTGGACAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_P40N2	-----													
Pain_P40N2_1	GCTCCAGGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTATCGTTGTAATTGCTGATCAACCGTATCTGAGCTAACG													
Pain_P40N2_2	GCTCCAGGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTATCGTTGTAATTGCTGATCAACCGTATCTGAGCTAACG													
Consensus	GCTCCAGGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTATCGTTGTAATTGCTGATCAACCGTATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_P40N2	-----													
Pain_P40N2_1	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCGGTGTTGG													
Pain_P40N2_2	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCGGTGTTGG													
Consensus	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGTCGAGCTGAGACTCACTTCTGTGCTGATCAAACTAGATCCTCAGTGGCTCCGGGAGTTGCTAAACAAGTTTATGGTAGTTCAGTACCGGTGTTGG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_P40N2	-----													
Pain_P40N2_1	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCAGACTCTT													
Pain_P40N2_2	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCAGACTCTT													
Consensus	ACGGTGAAAAACATTTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAGGAGGAGAGAACAGTCATAACATCGCGAATTTACCCAACAAAGGCAGTGAATGGAGCAGCAGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_P40N2	-----													
Pain_P40N2_1	CGTTTTCAACAAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGAGATTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Pain_P40N2_2	CGTTTTCAACAAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGAGATTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													
Consensus	CGTTTTCAACAAATGCCACAGGGGCTAGCGTGACTGCCTCCGTCAGAGATTGGTCACTTGAGTCGGCTAATATTTCGATCCTTCCCCTTGCAAGACTTGTAAC													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P54N	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
Pain_P54N_2	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
Pain_P54N_1	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
Pain_P54N_4	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
Pain_P54N_5	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
Consensus	AT	G	G	C	C	A	G	T	A	C	C	A	T	T
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P54N	T	C	T	T	T	G	C	T	T	T	C	G	A	T
Pain_P54N_2	T	C	T	T	T	G	C	T	T	T	C	G	A	T
Pain_P54N_1	T	C	T	T	T	G	C	T	T	T	C	G	A	T
Pain_P54N_4	T	C	T	T	T	G	C	T	T	T	C	G	A	T
Pain_P54N_5	T	C	T	T	T	G	C	T	T	T	C	G	A	T
Consensus	T	C	T	T	T	G	C	T	T	T	C	G	A	T
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P54N	T	G	T	C	G	T	A	G	T	A	C	A	G	T
Pain_P54N_2	T	G	T	C	G	T	A	G	T	A	C	A	G	T
Pain_P54N_1	T	G	T	C	G	T	A	G	T	A	C	A	G	T
Pain_P54N_4	T	G	T	C	G	T	A	G	T	A	C	A	G	T
Pain_P54N_5	T	G	T	C	G	T	A	G	T	A	C	A	G	T
Consensus	T	G	T	C	G	T	A	G	T	A	C	A	G	T
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P54N	T	G	G	T	A	T	C	A	A	T	A	T	C	A
Pain_P54N_2	T	G	G	T	A	T	C	A	A	T	A	T	C	A
Pain_P54N_1	T	G	G	T	A	T	C	A	A	T	A	T	C	A
Pain_P54N_4	T	G	G	T	A	T	C	A	A	T	A	T	C	A
Pain_P54N_5	T	G	G	T	A	T	C	A	A	T	A	T	C	A
Consensus	T	G	G	T	A	T	C	A	A	T	A	T	C	A
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_P54N	A	C	G	A	T	A	A	A	C	G	G	T	G	T
Pain_P54N_2	A	C	G	A	T	A	A	A	C	G	G	T	G	T
Pain_P54N_1	A	C	G	A	T	A	A	A	C	G	G	T	G	T
Pain_P54N_4	A	C	G	A	T	A	A	A	C	G	G	T	G	T
Pain_P54N_5	A	C	G	A	T	A	A	A	C	G	G	T	G	T
Consensus	A	C	G	A	T	A	A	A	C	G	G	T	G	T
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_P54N	T	C	C	T	C	T	T	C	T	A	G	A	C	T
Pain_P54N_2	T	C	C	T	C	T	T	C	T	A	G	A	C	T
Pain_P54N_1	T	C	C	T	C	T	T	C	T	A	G	A	C	T
Pain_P54N_4	T	C	C	T	C	T	T	C	T	A	G	A	C	T
Pain_P54N_5	T	C	C	T	C	T	T	C	T	A	G	A	C	T
Consensus	T	C	C	T	C	T	T	C	T	A	G	A	C	T
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_P54N	A	C	A	A	T	C	G	G	T	C	A	G	A	T
Pain_P54N_2	A	C	A	A	T	C	G	G	T	C	A	G	A	T
Pain_P54N_1	A	C	A	A	T	C	G	G	T	C	A	G	A	T
Pain_P54N_4	A	C	A	A	T	C	G	G	T	C	A	G	A	T
Pain_P54N_5	A	C	A	A	T	C	G	G	T	C	A	G	A	T
Consensus	A	C	A	A	T	C	G	G	T	C	A	G	A	T
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_P54N	T	T	A	C	C	G	G	T	A	C	A	A	A	A
Pain_P54N_2	T	T	A	C	C	G	G	T	A	C	A	A	A	A
Pain_P54N_1	T	T	A	C	C	G	G	T	A	C	A	A	A	A
Pain_P54N_4	T	T	A	C	C	G	G	T	A	C	A	A	A	A
Pain_P54N_5	T	T	A	C	C	G	G	T	A	C	A	A	A	A
Consensus	T	T	A	C	C	G	G	T	A	C	A	A	A	A
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_P54N	G	A	C	A	A	G	A	A	C	A	A	A	A	A
Pain_P54N_2	G	A	C	A	A	G	A	A	C	A	A	A	A	A
Pain_P54N_1	G	A	C	A	A	G	A	A	C	A	A	A	A	A
Pain_P54N_4	G	A	C	A	A	G	A	A	C	A	A	A	A	A
Pain_P54N_5	G	A	C	A	A	G	A	A	C	A	A	A	A	A
Consensus	G	A	C	A	A	G	A	A	C	A	A	A	A	A
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_P54N	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_P54N_2	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_P54N_1	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_P54N_4	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Pain_P54N_5	T	G	G	A	T	T	G	A	T	T	G	A	T	T
Consensus	T	G	G	A	T	T	G	A	T	T	G	A	T	T
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_P54N	T	T	A	A	G	C	T	T	A	A	G	C	T	T
Pain_P54N_2	T	T	A	A	G	C	T	T	A	A	G	C	T	T
Pain_P54N_1	T	T	A	A	G	C	T	T	A	A	G	C	T	T
Pain_P54N_4	T	T	A	A	G	C	T	T	A	A	G	C	T	T
Pain_P54N_5	T	T	A	A	G	C	T	T	A	A	G	C	T	T
Consensus	T	T	A	A	G	C	T	T	A	A	G	C	T	T
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_P54N	G	C	T	C	A	G	G	A	A	T	T	G	A	T
Pain_P54N_2	G	C	T	C	A	G	G	A	A	T	T	G	A	T
Pain_P54N_1	G	C	T	C	A	G	G	A	A	T	T	G	A	T
Pain_P54N_4	G	C	T	C	A	G	G	A	A	T	T	G	A	T
Pain_P54N_5	G	C	T	C	A	G	G	A	A	T	T	G	A	T
Consensus	G	C	T	C	A	G	G	A	A	T	T	G	A	T
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_P54N	C	C	A	G	T	T	A	C	T	T	C	A	A	G
Pain_P54N_2	C	C	A	G	T	T	A	C	T	T	C	A	A	G
Pain_P54N_1	C	C	A	G	T	T	A	C	T	T	C	A	A	G
Pain_P54N_4	C	C	A	G	T	T	A	C	T	T	C	A	A	G
Pain_P54N_5	C	C	A	G	T	T	A	C	T	T	C	A	A	G
Consensus	C	C	A	G	T	T	A	C	T	T	C	A	A	G
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_P54N	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_P54N_2	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_P54N_1	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_P54N_4	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Pain_P54N_5	A	C	G	G	T	A	A	A	A	C	A	T	T	C
Consensus	A	C	G	G	T	A	A	A	A	C	A	T	T	C
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_P54N	C	G	T	T	T	C	A	C	A	A	T	T	G	A
Pain_P54N_2	C	G	T	T	T	C	A	C	A	A	T	T	G	A
Pain_P54N_1	C	G	T	T	T	C	A	C	A	A	T	T	G	A
Pain_P54N_4	C	G	T	T	T	C	A	C	A	A	T	T	G	A
Pain_P54N_5	C	G	T	T	T	C	A	C	A	A	T	T	G	A
Consensus	C	G	T	T	T	C	A	C	A	A	T	T	G	A

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_P40N1	ATGCCACGCGAGTACCACTCAGTTATGACCCGGAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACACGATTCGGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTC													
Pain_P40N2	ATGCCACGCGAGTACCACTCAGTTATGACCCGGAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACACGATTCGGGCCACCGGAATCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTC													
Pain_P18N2	ATGCCACGCGAGTACCATTCCAGTTATGACCTGGAAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCGGGCCACCGGAAGTCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Pain_P54N	ATGCCACGCGAGTACCATTCCAGTTATGACCTGGAAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACCTGATTCGGGCCACCGGAAGTCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Pain_P18N1	ATGCCACGCGAGTACCATTCCAGTTATGACCCGGAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCGGGCCACCGGAAGTCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTT													
Consensus	ATGCCACGCGAGTACCACTCAGTTATGACCCGGAARAACCTCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCGGGCCACCGGAAGTCCATTAAATCATCTCCGGCATTTTCTCTCTCTCTC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_P40N1	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTGCCATTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P40N2	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTGCCATTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P18N2	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTTCCCGTTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P54N	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTTCCCGTTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Pain_P18N1	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTTCCCGTTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTTTAGTCTTCTTTCCGATCTCTACACACCAATCACCAGGACTTGCAAGTAACTTCCCGTTTCGCCGGCGCCGCCGTCAGAGGTTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_P40N1	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
Pain_P40N2	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
Pain_P18N2	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
Pain_P54N	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
Pain_P18N1	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
Consensus	TGTCGTCAATGCTAGTCACTGTTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAGAAGTCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACACAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_P40N1	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
Pain_P40N2	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
Pain_P18N2	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
Pain_P54N	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
Pain_P18N1	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTATCAATCAATCCAGATTACGCTATTTGGGGAATATCACATGGGGCCATGCCGATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTGGCATGGTTCTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_P40N1	ACGATATTACCGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
Pain_P40N2	ACGATATTACCGGGGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
Pain_P18N2	ACGATATAACCGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
Pain_P54N	ACGATATAACCGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
Pain_P18N1	ACGATATTACCGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
Consensus	ACGATATTACCGGTGCTGGACTGGGTCCGCTACCATCTACCCGATGGTCAGATCATGATGCTTTATACCTGGTGACACTGATGATTATGTGCAAGTGCAAAATCTTGCGTACCCCGCAACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_P40N1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P40N2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P18N2	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P54N	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Pain_P18N1	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCAACCCGGTTCTGGTTCTCCACCCGGCATTGGTGTCAGAGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCARTGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_P40N1	ACAATCGGGTCTAGATTGGTAAACCGGGATTGCACTTGTTTATGAACCTTCAC													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_DA	ATGGCCACG	CAGTACC	ATTCCAG	TTATGACC	CGGAA	AACTCCG	CTCCATT	ACACATT	CTCCCGG	ATCAACC	CGATTCC	GGCCACC	GGAGTCC	CTTAAAT
Pain_SA	ATGGCCACG	CAGTACC	ATTCCAG	TTATGACC	CGGAA	AACTCCG	CTCCATT	ACACATT	CTCCCGG	ATCAACC	CGATTCC	GGCCACC	GGAGTCC	CTTAAAT
Pain_TN2	ATGGCCACG	CAGTACC	ATTCCAG	TTATGACC	CGGAA	AACTCCG	CTCCATT	ACACATT	CTCCCGG	ATCAACC	CGATTCC	GGCCACC	GGAGTCC	CTTAAAT
Pain_TN1	ATGGCCACG	CAGTACC	ATTCCAG	TTATGACC	CGGAA	AACTCCG	CTCCATT	ACACATT	CTCCCGG	ATCAACC	CGATTCC	GGCCACC	GGAGTCC	CTTAAAT
Consensus	ATGGCCACG	CAGTACC	ATTCCAG	TTATGACC	CGGAA	AACTCCG	CTCCATT	ACACATT	CTCCCGG	ATCAACC	CGATTCC	GGCCACC	GGAGTCC	CTTAAAT
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_DA	TCCTTTT	GCTTT	CTGTAG	CCTCT	TTCCG	ATCCT	CAACA	CCAG	TCACCG	GACTT	GCAG	AGTAA	CTCC	GAG
Pain_SA	TCCTTTT	GCTTT	CTGTAG	CCTCT	TTCCG	ATCCT	CAACA	CCAG	TCACCG	GACTT	GCAG	AGTAA	CTCC	GAG
Pain_TN2	TCCTTTT	GCTTT	CTGTAG	CCTCT	TTCCG	ATCCT	CAACA	CCAG	TCACCG	GACTT	GCAG	AGTAA	CTCC	GAG
Pain_TN1	TCCTTTT	GCTTT	CTGTAG	CCTCT	TTCCG	ATCCT	CAACA	CCAG	TCACCG	GACTT	GCAG	AGTAA	CTCC	GAG
Consensus	TCCTTTT	GCTTT	CTGTAG	CCTCT	TTCCG	ATCCT	CAACA	CCAG	TCACCG	GACTT	GCAG	AGTAA	CTCC	GAG
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_DA	TGTCGT	CAATG	CTAGT	CAC	ATTTCT	TATG	CGTGG	TCCA	ATGCT	ATGCT	TAGCT	GGCA	AGAA	CTGCT
Pain_SA	TGTCGT	CAATG	CTAGT	CAC	ATTTCT	TATG	CGTGG	TCCA	ATGCT	ATGCT	TAGCT	GGCA	AGAA	CTGCT
Pain_TN2	TGTCGT	CAATG	CTAGT	CAC	ATTTCT	TATG	CGTGG	TCCA	ATGCT	ATGCT	TAGCT	GGCA	AGAA	CTGCT
Pain_TN1	TGTCGT	CAATG	CTAGT	CAC	ATTTCT	TATG	CGTGG	TCCA	ATGCT	ATGCT	TAGCT	GGCA	AGAA	CTGCT
Consensus	TGTCGT	CAATG	CTAGT	CAC	ATTTCT	TATG	CGTGG	TCCA	ATGCT	ATGCT	TAGCT	GGCA	AGAA	CTGCT
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_DA	TGGTAT	CATCT	TTTTT	TATCA	ATA	CAATC	CA	GATTC	AGCT	ATTTG	GGGA	AAAT	ATC	ACAT
Pain_SA	TGGTAT	CATCT	TTTTT	TATCA	ATA	CAATC	CA	GATTC	AGCT	ATTTG	GGGA	AAAT	ATC	ACAT
Pain_TN2	TGGTAT	CATCT	TTTTT	TATCA	ATA	CAATC	CA	GATTC	AGCT	ATTTG	GGGA	AAAT	ATC	ACAT
Pain_TN1	TGGTAT	CATCT	TTTTT	TATCA	ATA	CAATC	CA	GATTC	AGCT	ATTTG	GGGA	AAAT	ATC	ACAT
Consensus	TGGTAT	CATCT	TTTTT	TATCA	ATA	CAATC	CA	GATTC	AGCT	ATTTG	GGGA	AAAT	ATC	ACAT
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_DA	ACGATAT	AACGG	TGCTG	GACTG	GGTCC	GCAC	CA	CTCCT	ACCCG	ATGGT	CAG	ATCAT	GATG	CTTT
Pain_SA	ACGATAT	AACGG	TGCTG	GACTG	GGTCC	GCAC	CA	CTCCT	ACCCG	ATGGT	CAG	ATCAT	GATG	CTTT
Pain_TN2	ACGATAT	AACGG	TGCTG	GACTG	GGTCC	GCAC	CA	CTCCT	ACCCG	ATGGT	CAG	ATCAT	GATG	CTTT
Pain_TN1	ACGATAT	AACGG	TGCTG	GACTG	GGTCC	GCAC	CA	CTCCT	ACCCG	ATGGT	CAG	ATCAT	GATG	CTTT
Consensus	ACGATAT	AACGG	TGCTG	GACTG	GGTCC	GCAC	CA	CTCCT	ACCCG	ATGGT	CAG	ATCAT	GATG	CTTT
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_DA	TCCTCT	CCTT	CTAG	ACTG	GGTCA	AGTAC	AA	GGCA	ACCCG	GTCTG	GTTCT	CCACC	CGG	ATTG
Pain_SA	TCCTCT	CCTT	CTAG	ACTG	GGTCA	AGTAC	AA	GGCA	ACCCG	GTCTG	GTTCT	CCACC	CGG	ATTG
Pain_TN2	TCCTCT	CCTT	CTAG	ACTG	GGTCA	AGTAC	AA	GGCA	ACCCG	GTCTG	GTTCT	CCACC	CGG	ATTG
Pain_TN1	TCCTCT	CCTT	CTAG	ACTG	GGTCA	AGTAC	AA	GGCA	ACCCG	GTCTG	GTTCT	CCACC	CGG	ATTG
Consensus	TCCTCT	CCTT	CTAG	ACTG	GGTCA	AGTAC	AA	GGCA	ACCCG	GTCTG	GTTCT	CCACC	CGG	ATTG
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_DA	ACAATC	GGGT	CTA	AGATT	GGA	TAA	ACG	GGT	ATTG	CG	CTTGT	TATG	AA	CTT
Pain_SA	ACAATC	GGGT	CTA	AGATT	GGA	TAA	ACG	GGT	ATTG	CG	CTTGT	TATG	AA	CTT
Pain_TN2	ACAATC	GGGT	CTA	AGATT	GGA	TAA	ACG	GGT	ATTG	CG	CTTGT	TATG	AA	CTT
Pain_TN1	ACAATC	GGGT	CTA	AGATT	GGA	TAA	ACG	GGT	ATTG	CG	CTTGT	TATG	AA	CTT
Consensus	ACAATC	GGGT	CTA	AGATT	GGA	TAA	ACG	GGT	ATTG	CG	CTTGT	TATG	AA	CTT
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_DA	TTTACC	CGGT	ATC	GACT	GAAAA	ACA	ACG	GGT	TGG	AC	ACAT	CAT	AT	A
Pain_SA	TTTACC	CGGT	ATC	GACT	GAAAA	ACA	ACG	GGT	TGG	AC	ACAT	CAT	AT	A
Pain_TN2	TTTACC	CGGT	ATC	GACT	GAAAA	ACA	ACG	GGT	TGG	AC	ACAT	CAT	AT	A
Pain_TN1	TTTACC	CGGT	ATC	GACT	GAAAA	ACA	ACG	GGT	TGG	AC	ACAT	CAT	AT	A
Consensus	TTTACC	CGGT	ATC	GACT	GAAAA	ACA	ACG	GGT	TGG	AC	ACAT	CAT	AT	A
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_DA	GACA	AGA	ACA	AA	TGG	AC	ACCC	GAT	AAC	CCG	GA	ATTG	G	GA
Pain_SA	GACA	AGA	ACA	AA	TGG	AC	ACCC	GAT	AAC	CCG	GA	ATTG	G	GA
Pain_TN2	GACA	AGA	ACA	AA	TGG	AC	ACCC	GAT	AAC	CCG	GA	ATTG	G	GA
Pain_TN1	GACA	AGA	ACA	AA	TGG	AC	ACCC	GAT	AAC	CCG	GA	ATTG	G	GA
Consensus	GACA	AGA	ACA	AA	TGG	AC	ACCC	GAT	AAC	CCG	GA	ATTG	G	GA
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_DA	TGGATT	GGGG	AA	ACTG	ATG	TGA	ATCTG	CTG	AC	TG	CAG	AGG	GAT	GG
Pain_SA	TGGATT	GGGG	AA	ACTG	ATG	TGA	ATCTG	CTG	AC	TG	CAG	AGG	GAT	GG
Pain_TN2	TGGATT	GGGG	AA	ACTG	ATG	TGA	ATCTG	CTG	AC	TG	CAG	AGG	GAT	GG
Pain_TN1	TGGATT	GGGG	AA	ACTG	ATG	TGA	ATCTG	CTG	AC	TG	CAG	AGG	GAT	GG
Consensus	TGGATT	GGGG	AA	ACTG	ATG	TGA	ATCTG	CTG	AC	TG	CAG	AGG	GAT	GG
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_DA	TTGA	AGCT	TAA	GAGT	GGGT	GAT	CCT	ATTG	T	AG	CA	AGT	CA	AT
Pain_SA	TTGA	AGCT	TAA	GAGT	GGGT	GAT	CCT	ATTG	T	AG	CA	AGT	CA	AT
Pain_TN2	TTGA	AGCT	TAA	GAGT	GGGT	GAT	CCT	ATTG	T	AG	CA	AGT	CA	AT
Pain_TN1	TTGA	AGCT	TAA	GAGT	GGGT	GAT	CCT	ATTG	T	AG	CA	AGT	CA	AT
Consensus	TTGA	AGCT	TAA	GAGT	GGGT	GAT	CCT	ATTG	T	AG	CA	AGT	CA	AT
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_DA	GCTC	CAGG	GA	TA	ATTG	AG	CA	GA	T	CA	TG	AGG	T	T
Pain_SA	GCTC	CAGG	GA	TA	ATTG	AG	CA	GA	T	CA	TG	AGG	T	T
Pain_TN2	GCTC	CAGG	GA	TA	ATTG	AG	CA	GA	T	CA	TG	AGG	T	T
Pain_TN1	GCTC	CAGG	GA	TA	ATTG	AG	CA	GA	T	CA	TG	AGG	T	T
Consensus	GCTC	CAGG	GA	TA	ATTG	AG	CA	GA	T	CA	TG	AGG	T	T
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_DA	CCAG	TTT	ACT	TTCT	AC	ATTT	CTA	AGG	AG	CTG	ATGG	T	CG	T
Pain_SA	CCAG	TTT	ACT	TTCT	AC	ATTT	CTA	AGG	AG	CTG	ATGG	T	CG	T
Pain_TN2	CCAG	TTT	ACT	TTCT	AC	ATTT	CTA	AGG	AG	CTG	ATGG	T	CG	T
Pain_TN1	CCAG	TTT	ACT	TTCT	AC	ATTT	CTA	AGG	AG	CTG	ATGG	T	CG	T
Consensus	CCAG	TTT	ACT	TTCT	AC	ATTT	CTA	AGG	AG	CTG	ATGG	T	CG	T
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_DA	ACGG	T	AAAA	AC	ATT	CG	AT	G	AG	T	ATT	TGG	T	G
Pain_SA	ACGG	T	AAAA	AC	ATT	CG	AT	G	AG	T	ATT	TGG	T	G
Pain_TN2	ACGG	T	AAAA	AC	ATT	CG	AT	G	AG	T	ATT	TGG	T	G
Pain_TN1	ACGG	T	AAAA	AC	ATT	CG	AT	G	AG	T	ATT	TGG	T	G
Consensus	ACGG	T	AAAA	AC	ATT	CG	AT	G	AG	T	ATT	TGG	T	G
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_DA	CGT	TTT	CA	AC	AA	TG	CC	AC	AG	GG	G	CT	AG	CT
Pain_SA	CGT	TTT	CA	AC	AA	TG	CC	AC	AG	GG	G	CT	AG	CT
Pain_TN2	CGT	TTT	CA	AC	AA	TG	CC	AC	AG	GG	G	CT	AG	CT
Pain_TN1	CGT	TTT	CA	AC	AA	TG	CC	AC	AG	GG	G	CT	AG	CT
Consensus	CGT	TTT	CA	AC	AA	TG	CC	AC	AG	GG	G	CT	AG	CT

		1	10	20	30	40	50	60	70	80	90	100	110	120	130
BC14	Pain_DA	ATGGCCACCCAGTACCATTCCAGTTATGACCTGGAAAACTCCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
Consensus		ATGGCCACCGAGTACCATTCCAGTTATGACC C GGAAAACTCCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
		ATGGCCAC C AGTACCATTCCAGTTATGACC C GGAAAACTCCGCCTCCCATTTACACATTCTCCCGGATCAACCCGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTTCTCTCCTCTT													
		131	140	150	160	170	180	190	200	210	220	230	240	250	260
BC14	Pain_DA	TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAA CA AT TC ACCGGACTTGCAGAGTAACCTCCGTTCCGCCGGCCGCCGCTCAGAGGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
Consensus		TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAA CA AT TC ACCGGACTTGCAGAGTAACCTCCGTTCCGCCGGCCGCCGCTCAGAGGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
		TCCTTTTGTCTTCTGTAGCCTTCTTTCCGATCCTCAACAA CA AT TC ACCGGACTTGCAGAGTAACCTCCGTTCCGCCGGCCGCCGCTCAGAGGGTGTTTCTCAGGGAGTCTCCGATAGACTTTTCGAGA													
		261	270	280	290	300	310	320	330	340	350	360	370	380	390
BC14	Pain_DA	TGTGCTCAATGCTAGTCAC G TTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAAAGAACTGCTTACCATT TC AACTCAAAAAAATTGGATGAAC G GTAA TT AACTTTCTTATTTGACTTTTCT													
Consensus		TGTGCTCAATGCTAGTCAC A TTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAAAGAACTGCTTACCATT TC AACTCAAAAAAATTGGATGAAC G GTAA TT AACTTTCTTATTTGACTTTTCT													
		TGTGCTCAATGCTAGTCAC A TTTCTTATGCGTGGTCCAATGCTATGCTTAGCTGGCAAAAGAACTGCTTACCATT TC AACTCAAAAAAATTGGATGAAC G													
		391	400	410	420	430	440	450	460	470	480	490	500	510	520
BC14	Pain_DA	GTAA TT TCCTATTATT TT GTATCTTAGA TT GA AAAA AATTATAA TA CTTATACCGTTTTTTTTTTCTTAATGATATTTATGGCTATTGATCTGTTGGGGTATCTTTTGGA TT CTGATTGGATGCTATT													
Consensus														
														
		521	530	540	550	560	570	580	590	600	610	620	630	640	650
BC14	Pain_DA	CTGCAG ATCCTAATGGT GAGTTCAAAGTTAATTATCATCACTATTTTCTGTTAGTATTTAATTAA TT ATATATCTTTAA CC ATGA AA ATA AA ACTTTAAAGGCAGTA AA ATTCTCTCATGAGGTAA TT ATGG													
Consensus	 ATCCTAATGGT													
														
		651	660	670	680	690	700	710	720	730	740	750	760	770	780
BC14	Pain_DA	TTTTAAT TT GAATTAAGCCTATAAGTGGCA AA CAATCCATG TAT GAGCA AA TCATTAA TT CGGGTATGTCATCTCGGTTAATCCTTTTACCTTTTACTCA AA AGGA AA CTATTACTCCG TC CA AA ATAAT TT G													
Consensus														
														
		781	790	800	810	820	830	840	850	860	870	880	890	900	910
BC14	Pain_DA	ATGTTTCACAT AA TC AA TG T GATGTT AA TTTTTTTTTCA AA TTTACCCTTGGTATATACCT AA TCC T ATATGATTATGCC AA TC T AA T ATGA AA GA AA ATC AA T TA CA GA TATTTAG TC AC													
Consensus														
														
		911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
BC14	Pain_DA	AATTAATTCATGTTAA AA TATCA AA TAA TT TGGATTGGAGGTAGTACTAATTAGGA AA ATAATTAA GT TAATCATTTCAC TA AA CA TTGTTTAGACTAAGGATGA AA TAGGGGAGG AA TCA AA TATCT													
Consensus														
														
		1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
BC14	Pain_DA	TATTTTGTAA AT GGATAAGTATTTTGA AA TAA CA AA TT TTAGAA AA CA CG CAAGTCA AA TAGAGTAGGATTGATGGAGTGATTCTA AC CTTTCTAGATATTCATA AA AAT TT GGTGA AT TTTTTT													
Consensus														
														
		1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
BC14	Pain_DA	AATA AA CA CG ACAGT TT GATGGATTAGGCTTGTTGTTCCA AT TATA AT TGGGATTAA CA TGAGATCGTG GC AGCA AA GGTTTTTG GT TTTGGGTA AT TTTCCA AT AA AA AAT TA ACACATGATTGG CC CA													
Consensus														
														
		1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
BC14	Pain_DA	GTTTTATACAGT TT GGAA TC AA TC ACGTTATG GG TCAT CT TTTTTGTAGTA AT GTAA TA AT CC ATTAGTTGGCCCCC AT CCA AA TATTTGTCCATCTTCCAC TT GGTCATTTTCTCTTCTT													
Consensus														
														
		1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
BC14	Pain_DA	TTATTTTTTGA AA TGGAGTAGGTTATCTTGTGCCGCTTAGAG CA T TA CTATTACCATT TC GAGTCA AA AAAA AA TCA AT ATATATTATAAGGATA AA AATATATACAT AA ATTT CA TGAGTTAT													
Consensus														
														
		1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
BC14	Pain_DA	TTATTTAA AT TTTAGGGGAGGGAGC ATA CA TA GT AA CA TA CTAGTAA AT TGTTTAA GT AGTAGCTTGTTGA AG ATA AT CTTA AT TATTTCA AG GTCA AA AA TA TA AC TATG GC GA AA TTT TC AA													
Consensus														
														
		1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
BC14	Pain_DA	TAA AG AC CT TATCTTTTTG CC GA AA AA AG CA TAG CA AT TTTGGTAC GG AA CA TATTGAGAT TC TGTA AG ATTTTAC AA TTCA AA TTGCATGA AA AGTCTTAC CT AA TC CA AG TA AA ATACAT AT													
Consensus														
														
		1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920	1930	1940	1950
BC14	Pain_DA	TACTTTGA AA TTTCTACT AA CGTGA AA TA AT TGGTCA AC AGGT CC ATTGTACCACAGGGATGGTATCATCTTTTTTATCA AT AC AA TC CA GATTCAGCTATTTGGGGAA AT ATCA TA TAGGGCCAT GC													
Consensus	 CC ATTGTACCACAGGGATGGTATCATCTTTTTTATCA AT AC AA TC CA GATTCAGCTATTTGGGGAA AT ATCA TA TAGGGCCAT GC													
	 CC ATTGTACCACAGGGATGGTATCATCTTTTTTATCA AT AC AA TC CA GATTCAGCTATTTGGGGAA AT ATCA TA TAGGGCCAT GC													
		1951	1960	1970	1980	1990	2000	2010	2020	2030	2040	2050	2060	2070	2080
BC14	Pain_DA	GTATCCA AG GACTTGATCC AC TGGCTCTACTTG CC TTTGG CA TGGTTCCTGATCA AT GGTACGATAT AA ACGGTGTCTGGACTGGGTC CG CTACCATCCTACCCGATGGTCA GA TGATGCTTTATA													
Consensus		GTATCCA AG GACTTGATCC AC TGGCTCTACTTG CC TTTGG CA TGGTTCCTGATCA AT GGTACGATAT AA ACGGTGTCTGGACTGGGTC CG C ACCATCCTACCCGATGGTCA GA TGATGCTTTATA													
		GTATCCA AG GACTTGATCC AC TGGCTCTACTTG CC TTTGG CA TGGTTCCTGATCA AT GGTACGATAT AA ACGGTGTCTGGACTGGGTC CG C ACCATCCTACCCGATGGTCA GA TGATGCTTTATA													
		2081	2090	2100	2110	2120	2130	2140	2150	2160	2170	2180	2190	2200	2210
BC14	Pain_DA	CCGGTGA CA CTGATGATTATGTAC AG TGC AA AA CT TTGG CA TCC CA CA CA CTTATCTGATCCTCTCCTTCTAGACTGGGTC AA GTAC AA AGGC AA CCGGTCTGGTTCCTCC AC CCGGCATTGGT GT													
Consensus		CCGGTGA CA CTGATGATTATGTAC AG TGC AA AA CT TTGG CA TCC CA CA CA CTTATCTGATCCTCTCCTTCTAGACTGGGTC AA GTAC AA AGGC AA CCGGTCTGGTTCCTCC AC CCGGCATTGGT GT													
		CCGGTGA CA CTGATGATTATGTAC AG TGC AA AA CT TTGG CA TCC CA CA CA CTTATCTGATCCTCTCCTTCTAGACTGGGTC AA GTAC AA AGGC AA CCGGTCTGGTTCCTCC AC CCGGCATTGGT GT													
		2211	2220	2230	2240	2250	2260	2270	2280	2290	2300	2310	2320	2330	2340
BC14	Pain_DA	CAAGGACTTTAGAGAC CC GACC ACT GTG													
Consensus		CAAGGACTTTAGAGAC CC GACC ACT GTG													
		CAAGGACTTTAGAGAC CC GACC ACT GTG													
		2341	2350	2360	2370	2380	2390	2400	2410	2420	2430	2440	2450	2460	2470
BC14	Pain_DA	AAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGGTGGACTTTTACCCGGTATCGACTGA AA AA CA AA AC GGGTTGGACACATCATATACGCGCCGGGTGA AA AGCATG													
Consensus		AAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGGTGGACTTTTACCCGGTATCGACTGA AA AA CA AA AC GGGTTGGACACATCATATACGCGCCGGGTGA AA AGCATG													
		AAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGGTGGACTTTTACCCGGTATCGACTGA AA AA CA AA AC GGGTTGGACACATCATATACGCGCCGGGTGA AA AGCATG													
		2471	2480	2490	2500	2510	2520	2530	2540	2550	2560	2570	2580	2590	2600
BC14	Pain_DA	TGTTAA AG CAAGTTTAGATGAC AA TAA GC AA GA TCA CT ATGCTATTGGGAGCTATGACTTGAC AA GA AA TAA AT GGAC AC CCGAT AA CCCGGA AT TGGATTGTGGA AT TGGGTTGA AG CTGGATTATGG													
Consensus		TGTTAA AG CAAGTTTAGATGAC AA TAA GC AA GA TCA CT ATGCTATTGGGAGCTATGACTTGAC AA GA AA C AA AT GGAC AC CCGAT AA CCCGGA AT TGGATTGTGGA AT TGGGTTGA AG CTGGATTATGG													
		TGTTAA AG CAAGTTTAGATGAC AA TAA GC AA GA TCA CT ATGCTATTGGGAGCTATGACTTGAC AA GA AA C AA AT GGAC AC CCGAT AA CCCGGA AT TGGATTGTGGA AT TGGGTTGA AG CTGGATTATGG													
		2601	2610	2620	2630	2640	2650	2660	2670	2680	2690	2700	2710	2720	2730
BC14	Pain_DA	GA AA TATTATG CA TCA AA AGACATTTTATG AC CCG AA GA AA CA CG AGAGT ACT GTGGGGATGGATTGGGGAA CT GATG T GA AT CTG CT G AC CTG CA GA GG AGTGGG CA T CT G T AC AG GTATGGACT													
Consensus		GA AA TATTATG CA TCA AA AGACATTTTATG AC CCG AA GA AA CA CG AGAGT ACT GTGGGGATGGATTGGGGAA CT GATG T GA AT CTG CT G AC CTG CA GA GG AGTGGG CA T CT G T AC AG													
		GA AA TATTATG CA TCA AA AGACATTTTATG AC CCG AA GA AA CA CG AGAGT ACT GTGGGGATGGATTGGGGAA CT GATG T GA AT CTG CT G AC CTG CA GA GG AGTGGG CA T CT G T AC AG													
		2731	2740	2750	2760	2770	2780	2790	2800	2810	2820	2830	2840	2850	2860
BC14	Pain_DA	TCGATTAA CA CATTGTTTGTGTTTAGTTTG CA CCATAC CA AA TA ATCGTGCGA AA TTATATCTATCAGTAGGGAA TT CTTATTTAG AA AA AA AGT T GTATATATCA AT GCATT T GTGGTGA AG TGA													
Consensus														
														
		2861	2870	2880	2890	2900	2910	2920	2930	2940	2950	2960	2970	2980	2990
BC14	Pain_DA	ATTGGAA TT TTTGTG TA AGTATAT CT CA AT TATATATGA AA CA TC TCTA AT AA TT TTG TA ATACGA AT ATA TA CA GA AGTAT TC CA AG GA CA GTG CT TTACG CA CA GA AGACAGGGACACATCTACT TC CA													
Consensus	 AG TAT TC CA AG GA CA GTG CT TTACG CA CA GA AGACAGGGACACATCTACT TC CA													
	 AG TAT TC CA AG GA CA GTG CT TTACG CA CA GA AGACAGGGACACATCTACT TC CA													
		2991	3000	3010	3020	3030	3040	3050	3060	3070	3080	3090	3100	3110	3120
BC14	Pain_DA	GTGGCCAGTTGA AG AA AT TGA AA GC TT AA AG CGGGTGATCCTATTGTTAA GC AA GT CA AT CTTCA AC CA GG TTCA AT TGAG CT ACTCCATGTTGACTCAG CT G CA GA GG TTGTTGTTGCGCGACTTTGTTT													
Consensus		GTGGCCAGTTGA AG AA AT TGA AA GC TT AA AG CGGGTGATCCTATTGTTAA GC AA GT CA AT CTTCA AC CA GG TTCA AT TGAG CT ACTCCATGTTGACTCAG CT G CA GA GG													
		GTGGCCAGTTGA AG AA AT TGA AA GC TT AA AG CGGGTGATCCTATTGTTAA GC AA GT CA AT CTTCA AC CA GG TTCA AT TGAG CT ACTCCATGTTGACTCAG CT G CA GA GG													
		3121	3130	3140	3150	3160	3170	3180	3190	3200	3210	3220	3230	3240	3250
BC14	Pain_DA	AAAATTACA AA CTTTAT ACT TATACGCGCTT AT CTG CA GTCTTAA AA CTTGT T GGCTATTG T GCAGTTGGATATAGA AG CTCATT T GA AG TGGACAA AG TCGCGCTCCAGGGAA TA ATTGA AG CA GA T													
Consensus	 TT GGATATAGA AG CTCATT T GA AG TGGACAA AG TCGCGCTCCAGGGAA TA ATTGA AG CA GA T													
	 TT GGATATAGA AG CTCATT T GA AG TGGACAA AG TCGCGCTCCAGGGAA TA ATTGA AG CA GA T													
		3251	3260	3270	3280	3290	3300	3310	3320	3330	3340	3350	3360	3370	3380
BC14	Pain_DA	CATGTAGGTTTCAGCTG CT CTACTAGTGGAGG TG CTG TAG CA GA GGC AT TTTGGGACCATT T GGTGT CG TGTA AT TG CT GATCA AA CGCTATCTGAGCTA AC CGCAG TT TACTTCTACATTTCTAA AG													
Consensus		CATGTAGGTTTCAGCTG CT CTACTAGTGGAGG TG CTG TAG CA GA GGC AT TTTGGGACCATT T GGTGT CG TGTA AT TG CT GATCA AA C CTATCTGAGCTA AC CGCAG TT TACTTCTACATTTCTAA AG													
		CATGTAGGTTTCAGCTG CT CTACTAGTGGAGG TG CTG TAG CA GA GGC AT TTTGGGACCATT T GGTGT CG TGTA AT TG CT GATCA AA C CTATCTGAGCTA AC CGCAG TT TACTTCTACATTTCTAA AG													
		3381	3390	3400	3410	3420	3430	3440	3450	3460	3470	3480	3490	3500	3510
BC14	Pain_DA	GAGCTGATGG C CGAGCTGAGACTCA CT TCTGTGCTGATCA AA CC AG GT T GCTTCTATTTTCTCTATCTGGCACAA TT AA TT TGTCAT AT AGT CT CTGTA AA ATGGAGATGGATA AA AGTAGCGCGTTA													
Consensus		GAGCTGATGG C CGAGCTGAGACTCA CT TCTGTGCTGATCA AA CC AG GT T GCTTCTATTTTCTCTATCTGGCACAA TT AA TT TGTCAT AT AGT CT CTGTA AA ATGGAGATGGATA AA AGTAGCGCGTTA													
		GAGCTGATGG C CGAGCTGAGACTCA CT TCTGTGCTGATCA AA CC AG GT T GCTTCTATTTTCTCTATCTGGCACAA TT AA TT TGTCAT AT AGT CT CTGTA AA ATGGAGATGGATA AA AGTAGCGCGTTA													
		3511	3520	3530	3540	3550	3560	3570	3580	3590	3600	3610	3620	3630	3640
BC14	Pain_DA	TGATCTGATATATG CAG AT CT CTCAGAGG CT CCGGGAG TG CTAA CA AG TT TATGGT AG TTCAGTACC CG TGTTGGACG GT GA AA AA CA CTTCGATGAGAT TT TGGT AG TAGTGATGATGAT TC CTTATTT													
Consensus	 AT CTCTCAGAGG CT CCGGGAG TG CTAA CA AG TT TATGGT AG TTCAGTACC CG TGTTGGACG GT GA AA AA CA CTTCGATGAGAT TT TGGT AG													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	ATGGCCACGCAGTACCATTCCAGTTATGACCTGGAAACTCCGCTCCCATTAACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
Pain_SN*	ATGGCCACGCAGTACCATTCCAGTTATGACCTGGAAACTCCGCTCCCATTAACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
Consensus	ATGGCCACGCAGTACCATTCCAGTTATGACCTGGAAACTCCGCTCCCATTAACACATTCTCTCCGGATCAACCTGATTCCGGCCACCGGAAGTCCCTTAAATCATCTCCGGCATTTCCTCTCCTCTT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Pain_SN*	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
Consensus	TCCTTTTGCTTTCTGTAGCCTTCTTTCCGATCCTCAACAACCAATCACCGGACTTGCAGAGTAACCTCCGTTTCGCCGGCGCCGCGTCAAGAGGTGTTTCTCAGGGAGTCTCCGATAAGACTTTTCGAGA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
Pain_SN*	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
Consensus	TGTCGTCATGCTAGTCACGTTTCTTATGCGTGGTCCATGCTATGCTTAGCTGGCAAGAACTGCTTACCATTTCACCTCAAAAAAATTGGATGAACGATCCTAATGGTCCATTGTACCACAGGGA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Pain_SN*	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
Consensus	TGGTATCATCTTTTTTATCAATACATCCAGATTCAGCTATTTGGGGAATATCACATGGGGCCATGCCGTATCCAGGACTTGATCCACTGGCTCTACTTGCCTTTTGCCATGGTTCTGATCAATGGT													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	ACGATATAAACGGTGTCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Pain_SN*	ACGATATAAACGGTGTCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
Consensus	ACGATATAAACGGTGTCTGGACTGGGTCCGCTACCATCCTACCCGATGGTCAGATCATGATGCTTTATACCGGTGACACTGATGATTATGTACAGTGCAAAATCTTGCGTACCCACCACTTATCTGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCACCCGGTCTGGTTCCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
Pain_SN*	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCACCCGGTCTGGTTCCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
Consensus	TCCTCTCCTTCTAGACTGGGTCAAGTACAAAGGCACCCGGTCTGGTTCCTCCACCCGGCATTGGTGTCAAGGACTTTAGAGACCCGACCACTGCTTGGACCGGACCCCAAAATGGGCAATGGCTTTTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACTTGTTTATGAACCTTCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Pain_SN*	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACTTGTTTATGAACCTTCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
Consensus	ACAATCGGGTCTAAGATTGGTAAACGGGTATTGCACTTGTTTATGAACCTTCAACTTCACAAGCTTTAAGCTATTGGATGAAGTGCTGCATGCGGTTCCGGGTACGGGTATGTGGGAGTGTGTGGACT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TTTACCCGGTATCGACTGAAAAACAAACGGGTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Pain_SN*	TTTACCCGGTATCGACTGAAAAACAAACGGGTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
Consensus	TTTACCCGGTATCGACTGAAAAACAAACGGGTGGACACATCATATACGGCCCGGGTGTAAAGCATGTGTTAAAGCAAGTTAGATGACAAATAGCAAGATCACTATGCTATTGGGACGTATGACTT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	GACAAGAAACAATGGACACCCGATACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
Pain_SN*	GACAAGAAACAATGGACACCCGATACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
Consensus	GACAAGAAACAATGGACACCCGATACCCGGAATTGGATTGTGGAATTGGGTTGAAGCTGGATTATGGGAATATTATGCATCAAGACATTTTATGACCCGAGAAACACGAGAGTACTGTGGGGA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Pain_SN*	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
Consensus	TGGATTGGGGAAACTGATAGTGAATCTGCTGACCTGCAGAGGGGATGGGCATCTGTACAGAGTATTCCAGGACAGTGCTTTACGACAGAGAGACAGGGACACATCTACTTCAGTGGCCAGTTGAAGAA													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	TTGAAGCTTAAGAGCGGGTATCCTATTGTTAAGCAGTCAATCTTCAACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCGC													
Pain_SN*	TTGAAGCTTAAGAGCGGGTATCCTATTGTTAAGCAGTCAATCTTCAACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCGC													
Consensus	TTGAAGCTTAAGAGCGGGTATCCTATTGTTAAGCAGTCAATCTTCAACAGGTTCAATTGAGCTACTCCATGTTGACTCAGCTGCAGAGTTGGATATAGAGCCTCATTTGAAGTGGACAAGTCGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
Pain_SN*	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
Consensus	GCTCCAGGGAATAATTGAAGCAGATCATGTAGGTTTCAGCTGCTCTACTAGTGGAGGTGCTGCTAGCAGAGGCATTTTGGGACCATTTGGTGTCTGTTGAATTGCTGATCAACGCTATCTGAGCTAACG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAGATCCTCAGAGGCTCCGGGAGTTGCTAACAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Pain_SN*	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAGATCCTCAGAGGCTCCGGGAGTTGCTAACAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
Consensus	CCAGTTTACTTCTACATTTCTAAGGAGCTGATGGCCGAGCTGAGACTCACTTCTGTGCTGATCAACCAGATCCTCAGAGGCTCCGGGAGTTGCTAACAAGTTTATGGTAGTTTCAGTACCCGTGTTGG													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATACATCGCGAATTTACCAACAAGGACAGTGAATGGAGCAGCACGACTCTT													
Pain_SN*	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATACATCGCGAATTTACCAACAAGGACAGTGAATGGAGCAGCACGACTCTT													
Consensus	ACGGTGAAAAACATTCGATGAGATTATTGGTGGACCACTCAATTGTGGAGAGCTTTGCTCAAGGAGGAGAACAGTCATACATCGCGAATTTACCAACAAGGACAGTGAATGGAGCAGCACGACTCTT													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920			
Pain_SN	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
Pain_SN**	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCCTCCGTCAAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAA													
Pain_SN*	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCCTCCGTCAAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAA													
Consensus	CGTTTTCAACAATGCCACAGGGTCTAGCGTGACTGCCCTCCGTCAAGATTTGGTCACTTGAGTCGGCTAATATTCAATCCTTCCCCTTGCAAGACTTGTAA													

Appendix 3.2

A sequence was defined as an allele when it was found twice in two independent PCRs. Additionally, the consensus sequence of all alleles found in one genotype was used for allele definition when variable sequence polymorphisms occurred.

In the following Table, *invGE* and *invGF* alleles are listed, which do not correspond to this definition because only one full-length sequence was obtained. Nevertheless, several independent PCR amplifications gave rise to sequences that after partial sequencing indicated the existence of the alleles as mentioned above, clearly implying their existence. These additional sequences exhibited frame shifts or other modifications (e.g. missing or modified start and stop codons) and, therefore, were not completely sequenced. Furthermore, pyrosequencing analysis was carried out for allele specific SNPs demonstrating the presence of the latter SNPs in the corresponding genotype at genomic level.

Table A3.2.1: Overview of fully and partially sequenced *invGE* and *invGF* alleles.

Gene	Full-length allele	Additional partial sequences	Pyrosequencing assay
<i>invGE</i>	<i>E_SN2</i>	No additional sequences	Section 3.2.2.1.1, Figure 3.2.24
	<i>E_DN2</i>	No additional sequences	Section 3.2.2.1.1, Figure 3.2.24
<i>invGF</i>	<i>F_DN1</i>	2	Section 3.2.2.1.3, Figure 3.2.26
	<i>F_TN2</i>	2	Section 3.2.2.1.3, Figure 3.2.26
	<i>F_P40N1</i>	2	Section 3.2.2.1.4, Figure 3.2.27

1 10 20 30 40 50 60 70 80 90 100 110 120 130
|-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----|
E_SA MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SA3 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SA2 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SA1 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN1 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN1_1 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN2 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN3 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN3_1 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
E_SN3_2 MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA
Consensus MELFMKSSSLWGLEIYLFCEFIYVLSNINQVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWIHLEPAIYPSKKFDKYGAWSGSA

131 140 150 160 170 180 190 200 210 220 230 240 250 260
|-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----|
E_SA TILPNNKPVILYTGYYDSHDTQVQNYAIPANLSDPFLRKWYKPNMNPLIIPDNSINKTKFRDPTTAWMGVDGVMRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SA3 TILPNNKPVILYTGYYDSHDTQVQNYAIPANLSDPFLRKWYKPNMNPLIIPDNSINKTKFRDPTTAWMGVDGVMRIYIGSHRKHRGMALLYRGRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SA2 TILPNNKPVIXYTGYYDSHDTQVQNYAIPANLSDPFLRKWYKPNMNPLIIPDNSINKTKFRDPTTAWMGVDGVMRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SA1 TILPNNKPVILYTGYYDSHDTQVQNYAIPANLSDPFLRKWYKPNMNPLIIPDNSINKTKFRDPTTAWMGVDGVMRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN1 TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN1_1 TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN2 TILPNNKPIILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRNHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN3 TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN3_1 TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
E_SN3_2 TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNMNPLIIPDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG
Consensus TILPNNKP!ILYTGYYDSHDSQVQNYAIPANLSDPFLRKW!KPNMNPLI!PDNSINKTKFRDPTTAWMGQDGLWRIYIGSHRKHRGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFPVSLNNTNG

261 270 280 290 300 310 320 330 340 350 360 370 380 390
|-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----|
E_SA LDASYRGK-NVKYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SA3 LDASYRGK-NVKYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SA2 LDASYRGK-NVKYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SA1 LDASYRGK-NVKYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN1 LDASYRGKKKKYKHYVLKNSLDVNRFEYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN1_1 LDASYRGKKKKYKHYVLKNSLDVNRFEYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN2 LDASYRGKKKKYKHYVLKNSLDVNRFEYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN3 LDASYRGK-NVKHYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN3_1 LDASYRGK-NVKHYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
E_SN3_2 LDASYRGK-NVKHYVLKNSLDVNRFDYTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI-
Consensus LDASYRGK,nVKHYVLKNSLDVNRf#YTTIGHYDTRKORYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGTNESDVLPPDEIKKGWAGIQAIIPRKVWLDHSGKQLIQMPIEELETLRKQKI,

391 400 410 420 430 440 450 460 470 480 490 500 510 520
|-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----|
E_SA QLNNKKLSKGEMFEVKGISASQSDIEVSFSFSSLNKAQEQDPNWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SA3 QLNNKKLSKGEMFEVKGISASQSDIEVSFSFSSLNKAQEQDPNWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SA2 QSNNKKLSKGEMFEVKGISASQSDIEVSFSFSSLNKAQEQDPNWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SA1 QLNNKKLSKGEMFEVKGISASQSDIEVSFSFSSLNKAQEQDPNWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN1 QLNNKKLSKGEMFEVKGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN1_1 QLNNKKLSKGEMFEVKGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN2 QLNNKKLSKGEMFEVKGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN3 QLNNKKLSKGEMFEVEGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATVASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN3_1 QLNNKKLSKGEMFEVEGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATVASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
E_SN3_2 QLNNKKLSKGEMFEVEGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATVASKNLEEYTPVFFRVFKAQKNYKYLHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK
Consensus QLNNKKLSKGEMFEVKGISASQADIEVSFSFSSLNKAQEQDPKWADLYAQDVCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYK!LHCSDARRSTHRQNEAMYKPSFAGYVDVLDYDMK

521 530 540 550 560 570 580 586
|-----+-----+-----+-----+-----+-----+-----|
E_SA KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SA3 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SA2 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SA1 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN1 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN1_1 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN2 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN3 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN3_1 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
E_SN3_2 KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH
Consensus KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSHDVPMKH

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_DA	MELFMKSSSLWGLEIYLFCLFIIVLSNINKVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DA2	MELFMKSSSLWGLEIYLFCLFIIVLSNINKVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DA1	MELFMKSSSLWGLEIYLFCLFIIVLSNINKVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DN1	MELFMKSSSLWGLEIYLFCLFIIVLSNINGVFAASHNIFDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DN1_3	MELFMKSSSLWGLEIYLFCLFIIVLSNINGVFAASHNIFDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DN1_2	MELFMKSSSLWGLEIYLFCLFIIVLSNINGVFAASHNIFDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DN1_1	MELFMKSSSLWGLEIYLFCLFIIVLSNINGVFAASHNIFDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
E_DN2	MELFMKSSSLWGLEIYLFCLFIIVLSNINGVFAASHNIFDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
Consensus	MELFMKSSSLWGLEIYLFCLFIIVLSNINgVFashN!FLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYVAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_DA	TILPNNKPVILYTGVDSDHTQVQNYAIPANLSDPFLRKWVKPNNNPLIIPDNSINKTKFRDPTTAWMGVDGVWRIVIGSMRKHRGMALLYRSRDFIKWVKAQHPLHSSPHTGNWECPDFFPVSLNNTNG													
E_DA2	TILPNNKPVILYTGVDSDHTQVQNYAIPANLSDPFLRKWVKPNNNPLIIPDNSINKTKFRDPTTAWMGVDGVWRIVIGSMRKHRGMALLYRSRDFIKWVKAQHPLHSSPHTGNWECPDFFPVSLNNTNG													
E_DA1	TILPNNKPVILYTGVDSDHTQVQNYAIPANLSDPFLRKWVKPNNNPLIIPDNSINKTKFRDPTTAWMGVDGVWRIVIGSMRKHRGMALLYRSRDFIKWVKAQHPLHSSPHTGNWECPDFFPVSLNNTNG													
E_DN1	TILPNNKPVILYTGVDSDHSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_DN1_3	TILPNNKPVILYTGVDSDHSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_DN1_2	TILPNNKPVILYTGVDSDHSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_DN1_1	TILPNNKPVILYTGVDSDHSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAWMGQDGLWRIVIGSMRKHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_DN2	TILPNNKPIILYTGVDSDHSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAWMGQDGLWRIVIGSMRNHRGMALLYRSRDFIKWTKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
Consensus	TILPNNKP!ILYTGVDSDHsQVQNYAIPANLSDPFLRKW!KPNNNPLI!PDNSINKTKFRDPTTAWMGqDGLWRIVIGSMRkHRGMALLYRSRDFIKW.KAQHPLHSSPHTGNWECPDFFPVSLkNTNG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_DA	LDASYRGK-NVKYVLKNSLDYNRFDYTTIGMYDTRKDRYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEELETLRKQKI-													
E_DA2	LDASYRGK-NVKYVLKNSLDYNRFDYTTIGMYDTRKDRYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEELETLRKQKI-													
E_DA1	LDASYRGK-NVKYVLKNSLDYNRFDYTTIGMYDTRKDRYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEELETLRKQKI-													
E_DN1	LDASYRGK-NVKYVLKNSLDYNRFEYTTIGMYDTKKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDPSGKQLIQWPIEELETLRKQKII													
E_DN1_3	LDASYRGK-NVKYVLKNSLDYNRFEYTTIGMYDTKKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDPSGKQLIQWPIEELETLRKQKII													
E_DN1_2	LDASYRGK-NVKYVLKNSLDYNRFEYTTIGMYDTKKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDPSGKQLIQWPIEELETLRKQKII													
E_DN1_1	LDASYRGK-NVKYVLKNSLDYNRFEYTTIGMYDTKKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDPSGKQLIQWPIEELETLRKQKII													
E_DN2	LDASYRGKKNVKHVLKNSLDYNRFEYTTIGMYDTKKDRYIPDNNSIDGSKGLRLDYGNFYASKSFYDLPMKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDPSGKQLIQWPIEELETLRKQKII													
Consensus	LDASYRGK.NVKyVLKNSLDYNRF#YTTIGMYDTkkDRYIPDNNSIDGsKGLRLDYGNFYASKSFYDpmKNRRIYWGHTNESDVLPDDEIKKGWAGIQAIIPRKVWLDpSGKQLIQWPIEELETLRKQKIi													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_DA	QLNNKKLSKGEMFEYKGISASQSDIEVSFSFSSLNKAQFDPNWADLYAQDYCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDMK													
E_DA2	QLNNKKLSKGEMFEYKGISASQSDIEVSFSFSSLNKAQFDPNWADLYAQDYCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDMK													
E_DA1	QLNNKKLSKGEMFEYKGISASQSDIEVSFSFSSLNKAQFDPNWADLYAQDYCAIKGSTIQGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDMK													
E_DN1	QLNNKKLSKGEMFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLVTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTK													
E_DN1_3	QLNNKKLSKGEMFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLVTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTK													
E_DN1_2	QLNNKKLSKGEMFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLVTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTK													
E_DN1_1	QLNNKKLSKGEMFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLVTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTK													
E_DN2	QLNNKKLSKGEMFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTK													
Consensus	QLNNKKLSKGEMFEYKGISASQaDIEVSFSFSSLNKAQFDPkWADLYAQDYCAIKGSTIQgGLGPFGLaTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYKPSFAGYVDYDLVDTk													
	521	530	540	550	560	570	580	586						
E_DA	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DA2	KLSLRGLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DA1	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DN1	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DN1_3	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DN1_2	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DN1_1	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
E_DN2	KLSLRSLIDNSYVESFGAGGKTCITSRVYPTLAIHNAHLFVFNNNGSETITITETLNAWSMDVPKMH													
Consensus	KLSLRsLIDNSYVESFGAGGKTCITSRVYPTLAIH#NAHLFVFNNNGSETITITETLNAWSMDVPKMH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_SA	-----													
E_SA2	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTGGACTTGCAATCTTCAA													
E_SA3	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTGGACTTGCAATCTTCAA													
E_SA1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTGGACTTGCAATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTGGACTTGCAATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_SA	-----													
E_SA2	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACATCCAA													
E_SA3	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACATCCAA													
E_SA1	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACATCCAA													
Consensus	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACATCCAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_SA	-----													
E_SA2	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTACCCATCCAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC													
E_SA3	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTACCCATCCAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC													
E_SA1	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCAGTCTCAAAAGGCTTGATAAATTGGATCCATTAGAACCCGCAATTTACCCATCCAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC													
Consensus	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCAGTCTCAAAAGaCTTGATAAATTGGATCCATTAGAACCCGCAATTTACCCATCCAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_SA	-----													
E_SA2	ACTATTCTACCTAATAACAAGCCTGTTATTATATACACTGGAGTAGTAGATTCTCATGTAATCTCAAGTTCAAACTATGCCATCCAGCTAATCTGTCTGATCCGTTTCTTCGTAATGGGTCAAAACCA													
E_SA3	ACTATTCTACCTAATAACAAGCCTGTTATTATATACACTGGAGTAGTAGATTCTCATGTAATCTCAAGTTCAAACTATGCCATCCAGCTAATCTGTCTGATCCGTTTCTTCGTAATGGGTCAAAACCA													
E_SA1	ACTATTCTACCTAATAACAAGCCTGTTATTATATACACTGGAGTAGTAGATTCTCATGTAATCTCAAGTTCAAACTATGCCATCCAGCTAATCTGTCTGATCCGTTTCTTCGTAATGGGTCAAAACCA													
Consensus	ACTATTCTACCTAATAACAAGCCTGTTATTATATACACTGGAGTAGTAGATTCTCATGTAATCTCAAGTTCAAACTATGCCATCCAGCTAATCTGTCTGATCCGTTTCTTCGTAATGGGTCAAAACCA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_SA	-----													
E_SA2	ATAACAACCCGTTGATTATTCTTGACAAATAGCATCAACAAACCAATTTCTGATCCACACACCGCTTGGATGGGTGTAGATGGGGTTGGAGGATTGTAAATAGGAAGTATGAGAAACATAGAGGGAT													
E_SA3	ATAACAACCCGTTGATTATTCTTGACAAATAGCATCAACAAACCAATTTCTGATCCACACACCGCTTGGATGGGTGTAGATGGGGTTGGAGGATTGTAAATAGGAAGTATGAGAAACATAGAGGGAT													
E_SA1	ATAACAACCCGTTGATTATTCTTGACAAATAGCATCAACAAACCAATTTCTGATCCACACACCGCTTGGATGGGTGTAGATGGGGTTGGAGGATTGTAAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTATTCTTGACAAATAGCATCAACAAACCAATTTCTGATCCACACACCGCTTGGATGGGTGTAGATGGGGTTGGAGGATTGTAAATAGGAAGTATGAGAAACATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_SA	-----													
E_SA2	GGCATTATTGTATAGAGTAGAGACTTCATTAAATGGGTCAAGGCCAACACCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAATAATACTAATGGT													
E_SA3	GGCATTATTGTATAGAGTAGAGACTTCATTAAATGGGTCAAGGCCAACACCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAATAATACTAATGGT													
E_SA1	GGCATTATTGTATAGAGTAGAGACTTCATTAAATGGGTCAAGGCCAACACCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAATAATACTAATGGT													
Consensus	GGCATTATTGTATAGAGTAGAGACTTCATTAAATGGGTCAAGGCCAACACCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAATAATACTAATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_SA	-----													
E_SA2	TTAGATGCATCGTATCGCGGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATAACATT													
E_SA3	TTAGATGCATCGTATCGCGGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATAACATT													
E_SA1	TTAGATGCATCGTATCGCGGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATAACATT													
Consensus	TTAGATGCATCGTATCGCGGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATAACATT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_SA	-----													
E_SA2	CTATCGATGGTTGTAGGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAAATTTGTGGGGTTGGACCAATGAATCAGATGTTTACCTGACGA													
E_SA3	CTATCGATGGTTGTAGGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAAATTTGTGGGGTTGGACCAATGAATCAGATGTTTACCTGACGA													
E_SA1	CTATCGATGGTTGTAGGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAAATTTGTGGGGTTGGACCAATGAATCAGATGTTTACCTGACGA													
Consensus	CTATCGATGGTTGTAGGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAAATTTGTGGGGTTGGACCAATGAATCAGATGTTTACCTGACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_SA	-----													
E_SA2	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGAAAGATATGGCTCGATCATAGTGGTAACAAATTGATTCAATGGCCTATTGAAGAATTAGAACCCTAAGAAACAAAGATCCAATTG													
E_SA3	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_SN1	-----													
E_SN1_1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAATTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAATTATTTATTTTGCCTCTTTA A AGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_SN1	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_SN1_1	-----													
E_SN1_1	GTGCTATTAGTGTCAAGAATGTTTCATAGAAGCTGGTTTTCAATTTCAACCTCCTAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAA													
Consensus	GTGCTATTAGTGTCAAGAATGTTTCATAGAAGCTGGTTTTCAATTTCAACCTCCTAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAA													
E_SN1	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_SN1_1	-----													
E_SN1_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
E_SN1	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_SN1_1	-----													
E_SN1_1	ACTATTCTACCAATAACAAGCCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
Consensus	ACTATTCTACCAATAACAAGCCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
E_SN1	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_SN1_1	-----													
E_SN1_1	ATAACAACCCGTTGATTGTTCTTGACAATAGCATCAACAAACCAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTTCTTGACAATAGCATCAACAAACCAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_SN1	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_SN1_1	-----													
E_SN1_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTATCTCCTCATACTGGAATTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTATCTCCTCATACTGGAATTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_SN1	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_SN1_1	-----													
E_SN1_1	TTAGATGCATCGTATCGCGGAAAAAAGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACA													
Consensus	TTAGATGCATCGTATCGCGGAAAAAAGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACA													
E_SN1	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_SN1_1	-----													
E_SN1_1	ATTCTATCGATGGTTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGA													
Consensus	ATTCTATCGATGGTTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGA													
E_SN1	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_SN1_1	-----													
E_SN1_1	CGATGAATTAAAGAAAGGATGGGCTGGTATTCAAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACCCTTGAGAAGCAAAGATTCAA													
Consensus	CGATGAATTAAAGAAAGGATGGGCTGGTATTCAAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACCCTTGAGAAGCAAAGATTCAA													
E_SN1	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_SN1_1	-----													
E_SN1_1	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAGGTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTTTTTCAAGTTTGACGAGGGCCGAACAATTTGATCCTAATGGG													
Consensus	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAGGTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTTTTTCAAGTTTGACGAGGGCCGAACAATTTGATCCTAATGGG													
E_SN1	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
E_SN1_1	-----													
E_SN1_1	CCGACCTTTATGCCAAGATGTTTGTCGCAATTAAGGGTTTCGACTATCCAGGTGGGCTTGGGGCATTGGGGCTTGCAGCATTAGCTTCTAAAACTTGGAAGAATACACACCTGTTTTCTTTCGAGTGTT													
Consensus	CCGACCTTTATGCCAAGATGTTTGTCGCAATTAAGGGTTTCGACTATCCAGGTGGGCTTGGGGCATTGGGGCTTGCAGCATTAGCTTCTAAAACTTGGAAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_SN1	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
E_SN1_1	-----													
E_SN1_1	TAAGGCTCAAAGAATTATAAGATTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGAGAG													
Consensus	TAAGGCTCAAAGAATTATAAGATTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGAGAG													
E_SN1	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
E_SN3	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTCATAGTTTTATCAAACATTAAATAGGGTGTGGCTTCTCATATAATTTTTTTGGACTTGCAATCTTCAA														
E_SN3_1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTATAGTTTTATCAAACATTAAATAGGGTGTGGCTTCTCATATAATTTTTTTGGACTTGCAATCTTCAA														
E_SN3_2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTATAGTTTTATCAAACATTAAATAGGGTGTGGCTTCTCATATAATTTTTTTGGACTTGCAATCTTCAA														
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTATAGTTTTATCAAACATTAAATAGGGTGTGGCTTCTCATATAATTTTTTTGGACTTGCAATCTTCAA														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
E_SN3	GTGCAATTAGTGTCAAGAAATGTTTCATAGAACTAGTTTTTCATTTTCACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
E_SN3_1	GTGCAATTAGTGTCAAGAAATGTTTCATAGAACTAGTTTTTCATTTTCACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
E_SN3_2	GTGCAATTAGTGTCAAGAAATGTTTCATAGAACTAGTTTTTCATTTTCACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
Consensus	GTGCAATTAGTGTCAAGAAATGTTTCATAGAACTAGTTTTTCATTTTCACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
E_SN3	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
E_SN3_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
E_SN3_2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
E_SN3	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
E_SN3_1	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
E_SN3_2	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
Consensus	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
E_SN3	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAAAATTTTCGTGATCCAACAACCGCTTGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
E_SN3_1	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAAAATTTTCGTGATCCAACAACCGCTTGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
E_SN3_2	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAAAATTTTCGTGATCCAACAACCGCTTGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
Consensus	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAAAATTTTCGTGATCCAACAACCGCTTGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
E_SN3	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC														
E_SN3_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC														
E_SN3_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC														
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
E_SN3	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
E_SN3_1	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
E_SN3_2	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
Consensus	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
E_SN3	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTTTATGACCCTATGAAGAATCGAAGAATTGTATGGGGTTGGACAAATGAATCAGATGTTTTACCTGACGA														
E_SN3_1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTTTATGACCCTATGAAGAATCGAAGAATTGTATGGGGTTGGACAAATGAATCAGATGTTTTACCTGACGA														
E_SN3_2	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTTTATGACCCTATGAAGAATCGAAGAATTGTATGGGGTTGGACAAATGAATCAGATGTTTTACCTGACGA														
Consensus	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTTTATGACCCTATGAAGAATCGAAGAATTGTATGGGGTTGGACAAATGAATCAGATGTTTTACCTGACGA														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
E_SN3	TGAATTAGAAGGATGGGCAGGAATTCAGCTATTCCACGTAAGTATGGCTCGACCCTAGTGGCAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
E_SN3_1	TGAATTAGAAGGATGGGCAGGAATTCAGCTATTCCACGTAAGTATGGCTCGACCCTAGTGGCAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
E_SN3_2	TGAATTAGAAGGATGGGCAGGAATTCAGCTATTCCACGTAAGTATGGCTCGACCCTAGTGGCAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
Consensus	TGAATTAGAAGGATGGGCAGGAATTCAGCTATTCCACGTAAGTATGGCTCGACCCTAGTGGCAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
E_SN3	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACAAAGGCCGAACAAATTTGATCCTAAATGGGCTG														
E_SN3_1	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACAAAGGCCGAACAAATTTGATCCTAAATGGGCTG														
E_SN3_2	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACAAAGGCCGAACAAATTTGATCCTAAATGGGCTG														
Consensus	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACAAAGGCCGAACAAATTTGATCCTAAATGGGCTG														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
E_SN3	ACCTTTATGCCAAGATGTTTGTCGTCATTAGGGGTTGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACAGTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
E_SN3_1	ACCTTTATGCCAAGATGTTTGTCGTCATTAGGGGTTGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACAGTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
E_SN3_2	ACCTTTATGCCAAGATGTTTGTCGTCATTAGGGGTTGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACAGTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
Consensus	ACCTTTATGCCAAGATGTTTGTCGTCATTAGGGGTTGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACAGTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
E_SN3	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
E_SN3_1	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
E_SN3_2	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
Consensus	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
E_SN3	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCACGTTAGCGATTCATAAATATGCACATTTATTTGCTTCAATAATG														
E_SN3_1	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCACGTTAGCGATTCATAAATATGCACATTTATTTGCTTCAATAATG														
E_SN3_2	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCACGTTAGCGATTCATAAATATGCACATTTATTTGCTTCAATAATG														
Consensus	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCACGTTAGCGATTCATAAATATGCACATTTATTTGCTTCAATAATG														
	1691	1700	1710	1720	1730	1740	1750	1755							
E_SN3	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
E_SN3_1	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
E_SN3_2	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
Consensus	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
E_DA	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTTGGACTTGCAATCTTCAA														
E_DA2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTTGGACTTGCAATCTTCAA														
E_DA1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTTGGACTTGCAATCTTCAA														
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTAAATAGGTGTTTGGTTCTCATATGTTTTTTTGGACTTGCAATCTTCAA														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
E_DA	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
E_DA2	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
E_DA1	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
Consensus	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
E_DA	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTACCCATCCAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC														
E_DA2	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTACCCATCCAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC														
E_DA1	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTACCCATCCAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC														
Consensus	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTACCCATCCAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCC														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
E_DA	ACTATTCTACCTAATAACAAGCCTGTTATTTTATACACTGGAGTAGTAGATTCTCATGATACTCAAGTTCAAAACCTATGCCATCCAGCTAACTTGCTGATCCGTTTCTTCGTAATGGGTCAAACCCA														
E_DA2	ACTATTCTACCTAATAACAAGCCTGTTATTTTATACACTGGAGTAGTAGATTCTCATGATACTCAAGTTCAAAACCTATGCCATCCAGCTAACTTGCTGATCCGTTTCTTCGTAATGGGTCAAACCCA														
E_DA1	ACTATTCTACCTAATAACAAGCCTGTTATTTTATACACTGGAGTAGTAGATTCTCATGATACTCAAGTTCAAAACCTATGCCATCCAGCTAACTTGCTGATCCGTTTCTTCGTAATGGGTCAAACCCA														
Consensus	ACTATTCTACCTAATAACAAGCCTGTTATTTTATACACTGGAGTAGTAGATTCTCATGATACTCAAGTTCAAAACCTATGCCATCCAGCTAACTTGCTGATCCGTTTCTTCGTAATGGGTCAAACCCA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
E_DA	ATAACAACCCGTTGATTATTCCTGACAAATAGCATCAACAARACCAATTTTCGTGATCCAACAACCGCTTGGATGGGTGTAGATGGGGTTTGGAGGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
E_DA2	ATAACAACCCGTTGATTATTCCTGACAAATAGCATCAACAARACCAATTTTCGTGATCCAACAACCGCTTGGATGGGTGTAGATGGGGTTTGGAGGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
E_DA1	ATAACAACCCGTTGATTATTCCTGACAAATAGCATCAACAARACCAATTTTCGTGATCCAACAACCGCTTGGATGGGTGTAGATGGGGTTTGGAGGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
Consensus	ATAACAACCCGTTGATTATTCCTGACAAATAGCATCAACAARACCAATTTTCGTGATCCAACAACCGCTTGGATGGGTGTAGATGGGGTTTGGAGGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
E_DA	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGTCAAAGCCCAACACCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAATAATACTAATGGT														
E_DA2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGTCAAAGCCCAACACCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAATAATACTAATGGT														
E_DA1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGTCAAAGCCCAACACCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAATAATACTAATGGT														
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGTCAAAGCCCAACACCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCCCTGTATCATTAAATAATACTAATGGT														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
E_DA	TTAGATGCATCGTATCGCGAAAAAATGTCAAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATACAATT														
E_DA2	TTAGATGCATCGTATCGCGAAAAAATGTCAAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATACAATT														
E_DA1	TTAGATGCATCGTATCGCGAAAAAATGTCAAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATACAATT														
Consensus	TTAGATGCATCGTATCGCGAAAAAATGTCAAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGATTACTACACTATTGGCATGTATGACACCAGAAAGATAGGTACATTCTTGATACAATT														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
E_DA	CTATCGATGGTTGTAAGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAATTGTGTGGGGTTGGACCAATGAATCAGATGTTTTACCTGACGA														
E_DA2	CTATCGATGGTTGTAAGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAATTGTGTGGGGTTGGACCAATGAATCAGATGTTTTACCTGACGA														
E_DA1	CTATCGATGGTTGTAAGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAATTGTGTGGGGTTGGACCAATGAATCAGATGTTTTACCTGACGA														
Consensus	CTATCGATGGTTGTAAGGGATTGAGGCTTGATTATGGGAATTTCTATGCATCTAAATCATTTTATGATCCTACTAAGAAATCGAAGAATTGTGTGGGGTTGGACCAATGAATCAGATGTTTTACCTGACGA														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
E_DA	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGAAAAGTATGGCTCGATCATAGTGGTAAACAATTGATTCAATGGCCTATTGAAGAATTAGAACTTAAAGAAACAAAGATCCAATTG														
E_DA2	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGAAAAGTATGGCTCGATCATAGTGGTAAACAATTGATTCAATGGCCTATTGAAGAATTAGAACTTAAAGAAACAAAGATCCAATTG														
E_DA1	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGAAAAGTATGGCTCGATCATAGTGGTAAACAATTGATTCAATGGCCTATTGAAGAATTAGAACTTAAAGAAACAAAGATCCAATTG														
Consensus	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGAAAAGTATGGCTCGATCATAGTGGTAAACAATTGATTCAATGGCCTATTGAAGAATTAGAACTTAAAGAAACAAAGATCCAATTG														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
E_DA	AATAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGTCTGATATTGAAGTGTCATTTTCTTTTTCAAGTTTGAACAAGCTGAACAATTTGATCCTAATTGGGCTG														
E_DA2	AATAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGTCTGATATTGAAGTGTCATTTTCTTTTTCAAGTTTGAACAAGCTGAACAATTTGATCCTAATTGGGCTG														
E_DA1	AATAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGTCTGATATTGAAGTGTCATTTTCTTTTTCAAGTTTGAACAAGCTGAACAATTTGATCCTAATTGGGCTG														
Consensus	AATAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGTCTGATATTGAAGTGTCATTTTCTTTTTCAAGTTTGAACAAGCTGAACAATTTGATCCTAATTGGGCTG														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
E_DA	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGGTGGGCTTGGACCATTGTTGGGCTTGCAACATTAGCTTCTAAAACTTGGAGAATATACACCTGTTTTCTTTCGAGTGTTTAA														
E_DA2	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGGTGGGCTTGGACCATTGTTGGGCTTGCAACATTAGCTTCTAAAACTTGGAGAATATACACCTGTTTTCTTTCGAGTGTTTAA														
E_DA1	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGGTGGGCTTGGACCATTGTTGGGCTTGCAACATTAGCTTCTAAAACTTGGAGAATATACACCTGTTTTCTTTCGAGTGTTTAA														
Consensus	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGGTGGGCTTGGACCATTGTTGGGCTTGCAACATTAGCTTCTAAAACTTGGAGAATATACACCTGTTTTCTTTCGAGTGTTTAA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
E_DA	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
E_DA2	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
E_DA1	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
Consensus	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAAGCCCTCATTTGCTGGATATGTAGATGTAGATTAGTAGACATGAAGAAGTTA														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
E_DA	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTGGTGCTGGAGGAAAACATGCATAACATCGAGGGTGTATCCAACATTAGCAATTCATGATAATGCACATTTATTTGCTTCAATAATG														
E_DA2	TCTCTTAGGGTTTGATTGATAACTCAGTAGTGGAGAGTTTGGTGCTGGAGGAAAACATGCATAACATCGAGGGTGTATCCAACATTAGCAATTCATGATAATGCACATTTATTTGCTTCAATAATG														
E_DA1	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTGGTGCTGGAGGAAAACATGCATAACATCGAGGGTGTATCCAACATTAGCAATTCATGATAATGCACATTTATTTGCTTCAATAATG														
Consensus	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTGGTGCTGGAGGAAAACATGCATAACATCGAGGGTGTATCCAACATTAGCAATTCATGATAATGCACATTTATTTGCTTCAATAATG														
	1691	1700	1710	1720	1730	1740	1750	1755							
E_DA	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
E_DA2	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
E_DA1	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
Consensus	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130

E_DN1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_DN1.1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_DN1.3	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_DN1.2	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260

E_DN1	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAAATCCAA													
E_DN1.1	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAAATCCAA													
E_DN1.3	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAAATCCAA													
E_DN1.2	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAAATCCAA													
Consensus	GTGCTATTAGTGTCAGGAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAAATCCAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390

E_DN1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_DN1.1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_DN1.3	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_DN1.2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520

E_DN1	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATTTATGCATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAAACTTA													
E_DN1.1	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATTTATGCATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAAACTTA													
E_DN1.3	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATTTATGCATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAAACTTA													
E_DN1.2	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATTTATGCATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAAACTTA													
Consensus	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATTTATGCATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAAACTTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650

E_DN1	ATAACACCCGTTGATTGTACCTGACAATAGCATCAACAAACCAATTTCTGTATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_DN1.1	ATAACACCCGTTGATTGTACCTGACAATAGCATCAACAAACCAATTTCTGTATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_DN1.3	ATAACACCCGTTGATTGTACCTGACAATAGCATCAACAAACCAATTTCTGTATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_DN1.2	ATAACACCCGTTGATTGTACCTGACAATAGCATCAACAAACCAATTTCTGTATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACACCCGTTGATTGTACCTGACAATAGCATCAACAAACCAATTTCTGTATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780

E_DN1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTCTGTATCATTAAAAAATACAATGGC													
E_DN1.1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTCTGTATCATTAAAAAATACAATGGC													
E_DN1.3	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTCTGTATCATTAAAAAATACAATGGC													
E_DN1.2	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTCTGTATCATTAAAAAATACAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTCTGTATCATTAAAAAATACAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910

E_DN1	TTAGATGCATCGTATCGCGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
E_DN1.1	TTAGATGCATCGTATCGCGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
E_DN1.3	TTAGATGCATCGTATCGCGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
E_DN1.2	TTAGATGCATCGTATCGCGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
Consensus	TTAGATGCATCGTATCGCGAAAAAATGTCAAATATGTCCTTAAGAAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040

E_DN1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTCTATGACCCTATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_DN1.1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTCTATGACCCTATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_DN1.3	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTCTATGACCCTATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_DN1.2	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTCTATGACCCTATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
Consensus	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAAATCATTCTATGACCCTATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170

E_DN1	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTTCCGCGTAAGTATGGCTCGACCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
E_DN1.1	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTTCCGCGTAAGTATGGCTCGACCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
E_DN1.3	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTTCCGCGTAAGTATGGCTCGACCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
E_DN1.2	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTTCCGCGTAAGTATGGCTCGACCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
Consensus	TGAATTAAAGAAGGATGGGCTGGAATTCAGCTATTTCCGCGTAAGTATGGCTCGACCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300

E_DN1	TTGAACACACAGAGTTGAGCAGGGGAGAATGTTTGAGGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTTCAAGTTTGACACAGGCCGACAAATTTGATCCTAATGGG													
E_DN1.1	TTGAACACACAGAGTTGAGCAGGGGAGAATGTTTGAGGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTTCAAGTTTGACACAGGCCGACAAATTTGATCCTAATGGG													
E_DN1.3	TTGAACACACAGAGTTGAGCAGGGGAGAATGTTTGAGGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTTCAAGTTTGACACAGGCCGACAAATTTGATCCTAATGGG													
E_DN1.2	TTGAACACACAGAGTTGAGCAGGGGAGAATGTTTGAGGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTTCAAGTTTGACACAGGCCGACAAATTTGATCCTAATGGG													
Consensus	TTGAACACACAGAGTTGAGCAGGGGAGAATGTTTGAGGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTTCAAGTTTGACACAGGCCGACAAATTTGATCCTAATGGG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430

E_DN1	CCGACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAACTTGGAGAATACACACCTGTTTTCTTTTCGAGTGT													
E_DN1.1	CCGACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAACTTGGAGAATACACACCTGTTTTCTTTTCGAGTGT													
E_DN1.3	CCGACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAACTTGGAGAATACACACCTGTTTTCTTTTCGAGTGT													
E_DN1.2	CCGACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAACTTGGAGAATACACACCTGTTTTCTTTTCGAGTGT													
Consensus	CCGACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAACTTGGAGAATACACACCTGTTTTCTTTTCGAGTGT													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560

E_DN1	TAAGGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_DN1.1	TAAGGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_DN1.3	TAAGGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_DN1.2	TAAGGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
Consensus	TAAGGCTCAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690

E_DN1	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGA AAAACATGCATAACATCGAGGGTGATCCAACGTTAGCGATTCACAATAATGCACATTTATTTGTCTTCATA													
E_DN1.1	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGA AAAACATGCATAACATCGAGGGTGATCCAACGTTAGCGATTCACAATAATGCACATTTATTTGTCTTCATA													
E_DN1.3	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGA AAAACATGCATAACATCGAGGGTGATCCAACGTTAGCGATTCACAATAATGCACATTTATTTGTCTTCATA													
E_DN1.2	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGA AAAACATGCATAACATCGAGGGTGATCCAACGTTAGCGATTCACAATAATGCACATTTATTTGTCTTCATA													
Consensus	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGA AAAACATGCATAACATCGAGGGTGATCCAACGTTAGCGATTCACAATAATGCACATTTATTTGTCTTCATA													
	1691	1700	1710	1720	1730	1740	1750	1758						

E_DN1	ATGGATCTGAGACAAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_DN1.1	ATGGATCTGAGACAAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_DN1.3	ATGGATCTGAGACAAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_DN1.2	ATGGATCTGAGACAAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	ATGGATCTGAGACAAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_TN2	-----													
E_TN2_1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_TN2	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_TN2_1	-----													
E_TN2_1	GTGCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTCAATTTCAACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAA													
Consensus	GTGCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTCAATTTCAACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAA													
E_TN2	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_TN2_1	-----													
E_TN2_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
E_TN2	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_TN2_1	-----													
E_TN2_1	ACTATTCTACCAATAACAAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
Consensus	ACTATTCTACCAATAACAAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
E_TN2	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_TN2_1	-----													
E_TN2_1	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCAACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCAACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_TN2	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_TN2_1	-----													
E_TN2_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTTCTGTATCATTAAAAAATACAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTTCTGTATCATTAAAAAATACAATGGC													
E_TN2	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_TN2_1	-----													
E_TN2_1	TTAGATGCATCGTATCGCGGAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
Consensus	TTAGATGCATCGTATCGCGGAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
E_TN2	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_TN2_1	-----													
E_TN2_1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAAGAATCGAAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
Consensus	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAAGAATCGAAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_TN2	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_TN2_1	-----													
E_TN2_1	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGGCAAAAGATTATTCAA													
Consensus	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCTAGTGGTAACAACCTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGGCAAAAGATTATTCAA													
E_TN2	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_TN2_1	-----													
E_TN2_1	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTCAAGTTTGACAAGGCCGAACAATTTGATCCTAATGGG													
Consensus	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTCAAGTTTGACAAGGCCGAACAATTTGATCCTAATGGG													
E_TN2	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
E_TN2_1	-----													
E_TN2_1	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAAGAATACACACCTGTTTTCTTTCGAGTGTT													
Consensus	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_TN2	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
E_TN2_1	-----													
E_TN2_1	TAAGGCTCAAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
Consensus	TAAGGCTCAAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_TN2	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
E_TN2_1	-----													
E_TN2_1	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGTATCCAACGTTAGCGATTCAATAATGCACATTTATTTGTCTTCAATA													
Consensus	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGTATCCAACGTTAGCGATTCAATAATGCACATTTATTTGTCTTCAATA													
E_TN2	1691	1700	1710	1720	1730	1740	1750	1758						
E_TN2_1	-----													
E_TN2_1	ATGGATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	ATGGATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	ATGGATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCCTCTTATAGTTTATCAAACATTAAATGGGGTGTGTTGCTTCTCATATATTTTTTTGGACTTGCATCTTCAA													
E_TN3_1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCCTCTTATAGTTTATCAAACATTAAATGGGGTGTGTTGCTTCTCATATATTTTTTTGGACTTGCATCTTCAA													
E_TN3_2	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCCTCTTATAGTTTATCAAACATTAAATGGGGTGTGTTGCTTCTCATATATTTTTTTGGACTTGCATCTTCAA													
E_TN3_3	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCCTCTTATAGTTTATCAAACATTAAATGGGGTGTGTTGCTTCTCATATATTTTTTTGGACTTGCATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCCTCTTATAGTTTATCAAACATTAAATGGGGTGTGTTGCTTCTCATATATTTTTTTGGACTTGCATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	GTCCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAAA													
E_TN3_1	GTCCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAAA													
E_TN3_2	GTCCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAAA													
E_TN3_3	GTCCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAAA													
Consensus	GTCCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
E_TN3_1	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
E_TN3_2	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
E_TN3_3	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
Consensus	AGGATCAGTATGGGGTAATATTGTTTGGGCTCATTCACTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTGAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	ACTATTCTACCAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGCTGATCCATTTCTTCGTAATGGATCAAACCTA													
E_TN3_1	ACTATTCTACCAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGCTGATCCATTTCTTCGTAATGGATCAAACCTA													
E_TN3_2	ACTATTCTACCAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGCTGATCCATTTCTTCGTAATGGATCAAACCTA													
E_TN3_3	ACTATTCTACCAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGCTGATCCATTTCTTCGTAATGGATCAAACCTA													
Consensus	ACTATTCTACCAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGCTGATCCATTTCTTCGTAATGGATCAAACCTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	ATAACAACCCGTTGATTGTTCTGCAATAGCATCAACAACCAAAATTTCTGACCCACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_TN3_1	ATAACAACCCGTTGATTGTTCTGCAATAGCATCAACAACCAAAATTTCTGACCCACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_TN3_2	ATAACAACCCGTTGATTGTTCTGCAATAGCATCAACAACCAAAATTTCTGACCCACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_TN3_3	ATAACAACCCGTTGATTGTTCTGCAATAGCATCAACAACCAAAATTTCTGACCCACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTTCTGCAATAGCATCAACAACCAAAATTTCTGACCCACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC													
E_TN3_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC													
E_TN3_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC													
E_TN3_3	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTTCTGTATCATTAAAAAATACTAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	TTAGATGCATCGTATCGCGAAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAGATAGGTACATTCTGTATAATAATT													
E_TN3_1	TTAGATGCATCGTATCGCGAAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAGATAGGTACATTCTGTATAATAATT													
E_TN3_2	TTAGATGCATCGTATCGCGAAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAGATAGGTACATTCTGTATAATAATT													
E_TN3_3	TTAGATGCATCGTATCGCGAAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAGATAGGTACATTCTGTATAATAATT													
Consensus	TTAGATGCATCGTATCGCGAAAAAATGTCAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAGATAGGTACATTCTGTATAATAATT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	CTATCGATGGTTTCAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_TN3_1	CTATCGATGGTTTCAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_TN3_2	CTATCGATGGTTTCAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_TN3_3	CTATCGATGGTTTCAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
Consensus	CTATCGATGGTTTCAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGCAAAAGATTCATTG													
E_TN3_1	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGCAAAAGATTCATTG													
E_TN3_2	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGCAAAAGATTCATTG													
E_TN3_3	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGCAAAAGATTCATTG													
Consensus	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAGAATTAGAACATTAGAAGCAAAAGATTCATTG													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	AACAGAAGAGGTTGAGCAAGGGAGAAATGTTGAAGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTTCAGTTTGAACAGGGCCGAACAATTTGATCCTAATGGGCCG													
E_TN3_1	AACAGAAGAGGTTGAGCAAGGGAGAAATGTTGAAGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTTCAGTTTGAACAGGGCCGAACAATTTGATCCTAATGGGCCG													
E_TN3_2	AACAGAAGAGGTTGAGCAAGGGAGAAATGTTGAAGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTTCAGTTTGAACAGGGCCGAACAATTTGATCCTAATGGGCCG													
E_TN3_3	AACAGAAGAGGTTGAGCAAGGGAGAAATGTTGAAGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTTCAGTTTGAACAGGGCCGAACAATTTGATCCTAATGGGCCG													
Consensus	AACAGAAGAGGTTGAGCAAGGGAGAAATGTTGAAGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTTCAGTTTGAACAGGGCCGAACAATTTGATCCTAATGGGCCG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCAAGTGTTTAA													
E_TN3_1	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCAAGTGTTTAA													
E_TN3_2	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCAAGTGTTTAA													
E_TN3_3	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCAAGTGTTTAA													
Consensus	ACCTTTATGCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCAAGTGTTTAA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	GGCTCAAAGATTATAGGTTCTCATGTGCTCAGATGCGAGAGAGCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGGAGTTA													
E_TN3_1	GGCTCAAAGGATTATAGGTTCTCATGTGCTCAGATGCGAGAGAGCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGGAGTTA													
E_TN3_2	GGCTCAAAGATTATAGGTTCTCATGTGCTCAGATGCGAGAGAGCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGGAGTTA													
E_TN3_3	GGCTCAAAGATTATAGGTTCTCATGTGCTCAGATGCGAGAGAGCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGGAGTTA													
Consensus	GGCTCAAAGATTATAGGTTCTCATGTGCTCAGATGCGAGAGAGCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAGGAGTTA													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
		-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	-----+	
E_TN3	TCTCTTAGGAGTTTGATTGTAACCTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATACATCGAGGGGTATCCACGTTAGCGATTACAATAATGCACATTTATTTGTCTTCAATAATG													
E_TN3_1	TCTCTTAGGAGTTTGATTGTAACCTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATACATCGAGGGGTATCCACGTTAGCGATTACAATAATGCACATTTATTTGTCTTCAATAATG													
E_TN3_2	TCTCTTAGGAGTTTGATTGTAACCTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATACATCGAGGGGTATCCACGTTAGCGATTACAATAATGCACATTTATTTGTCTTCAATAATG													
E_TN3_3	TCTCTTAGGAGTTTGATTGTAACCTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATACATCGAGGGGTATCCACGTTAGCGATTACAATAATGCACATTTATTTGTCTTCAATAATG													
Consensus	TCTCTTAGGAGTTTGATTGTAACCTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATACATCGAGGGGTATCCACGTTAGCGATTACAATAATGCACATTTATTTGTCTTCAATAATG													
	1691	1700	1710	1720	1730	1740	1750	1755						
		-----+	-----+	-----+	-----+	-----+	-----+	-----+						
E_TN3	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_TN3_1	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_TN3_2	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_TN3_3	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P18N1	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKH WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N1_1	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKH WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N1_2	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKH WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N1_3	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKH WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N2	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKY WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N2_1	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKY WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N2_3	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKY WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N2_2	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKY WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
E_P18N2_4	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKY WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAIYPSKKFDKYGAMSGSA													
Consensus	MELFMKSSSLWGLEIYLFCLFIYLSNINGVFASHNIFLDLQSSSAISYKNVHRTGFHFQPPKy WINDPNAPMYYNGVYHLFYQYNPKGSVMGNIWAHSYSKDLINWHLLEPAiYPSKKFDKYGAMSGSA													

	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P18N1	TILPNNKPIILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR NHRGMALLYRSRDFIKWTKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N1_1	TILPNNKPIILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR NHRGMALLYRSRDFIKWTKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N1_2	TILPNNKPIILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR NHRGMALLYRSRDFIKWTKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N1_3	TILPNNKPIILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR NHRGMALLYRSRDFIKWTKAQHPLHSSPHTGNWECPDFFPVSLRNTNG													
E_P18N2	TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N2_1	TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N2_3	TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N2_2	TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
E_P18N2_4	TILPNNKPVILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													
Consensus	TILPNNKP!ILYTGYYDSHDSQVQNYAIPANLSDPFLRKWIKPNNNPLIYPDNSINKTKFRDPTTAMHGQDGLWRIYIGSMR KHRGMALLYRSRDFIKWAKAQHPLHSSPHTGNWECPDFFPVSLKNTNG													

	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P18N1	LDASYRGKKNYKHV LKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N1_1	LDASYRGKKNYKHV LKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N1_2	LDASYRGKKNYKHV LKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N1_3	LDASYRGKKNYKHV LKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N2	LDASYRGK-NVKY VLKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N2_1	LDASYRGK-NVKY VLKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N2_3	LDASYRGK-NVKY VPKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N2_2	LDASYRGK-NVKY VLKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
E_P18N2_4	LDASYRGK-NVKY VLKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													
Consensus	LDASYRGK.NVKy VLKNSLDVNRFEYTTIGHYDTKKDRIYIPDNNSIDGSKGLRLDYGNFYASKSFYDPMKNRRIYWGHTNESDVL PDEIK BGWAGIQAIRKVMWLDPSGKQLIQWPIEELETLRKQKII													

	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P18N1	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N1_1	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N1_2	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N1_3	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVTR													
E_P18N2	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N2_1	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N2_3	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N2_2	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ NEAMYKPSFAGYVDVLDVDTK													
E_P18N2_4	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ TEAMYKPSFAGYVDVLDVDTK													
Consensus	QLNNKKLSKGENFEYKGISASQADIEVSFSFSSLNKAQFDPKWADLYAQDYCAIKGSTIQGGLGPFGLYTLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQ nEAMYKPSFAGYVDVLDVDTK													

	521	530	540	550	560	570	580	586
	-----+-----+-----+-----+-----+-----+-----+-----							
E_P18N1	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N1_1	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N1_2	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N1_3	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N2	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N2_1	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N2_3	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N2_2	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT ETLNAWSHDVPKMH							
E_P18N2_4	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT IGTLNAWSHDVPKMH							
Consensus	KLSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHNNNAHLFVFNNGETITIT etLNAWSHDVPKMH							

Sequence	1	10	20	30	40	50	60	70	80	90	100	110	120	130																																																																																																																		
E_P40N1	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	L	F	I	V	L	S	N	I	N	G	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	L	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S	V	S	K	D	L	I	N	W	I	H	L	E	P	A	I	Y	P	S	K	K	F	D	K	Y	G	A	W	S	G	S	A
E_P40N1_1	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	L	F	I	V	L	S	N	I	N	G	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	L	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S	V	S	K	D	L	I	N	W	I	H	L	E	P	A	I	Y	P	S	K	K	F	D	K	Y	G	A	W	S	G	S	A
E_P40N1_2	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	L	F	I	V	L	S	N	I	N	G	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	S	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S	V	S	K	D	L	I	N	W	I	H	L	E	P	A	I	Y	P	S	K	K	F	D	K	Y	G	A	W	S	G	S	A
E_P40N2	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	F	F	I	V	L	S	N	I	N	K	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	L	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S	V	S	K	D	L	I	N	W	I	H	L	E	P	A	I	Y	P	S	K	K	F	D	K	Y	G	A	W	S	G	S	A
E_P40N2_4	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	F	F	I	V	L	S	N	I	N	K	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	L	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S	V	S	K	D	L	I	N	W	I	H	L	E	P	A	I	Y	P	S	K	K	F	D	K	Y	G	A	W	S	G	S	A
E_P40N2_2	M	E	L	F	M	K	S	S	L	M	G	L	E	I	Y	F	C	F	F	I	V	L	S	N	I	N	K	V	F	A	S	H	N	I	F	L	D	L	Q	S	S	S	A	I	S	V	K	N	V	H	R	T	G	F	H	F	Q	P	P	K	H	W	I	N	D	P	N	A	P	M	Y	Y	N	G	V	Y	H	L	F	Y	Q	Y	N	P	K	G	S	V	M	G	N	I	V	W	A	H	S																															

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	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_P18N2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
E_P18N2_1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
E_P18N2_3	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
E_P18N2_4	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
E_P18N2_2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCCTCTTATAGTTTTATCAAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCATCTTC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_P18N2	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P18N2_1	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P18N2_3	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P18N2_4	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P18N2_2	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
Consensus	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCTTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_P18N2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
E_P18N2_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
E_P18N2_3	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
E_P18N2_4	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
E_P18N2_2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTGTCTCAAAAGACTTGATAAATTGGATCCATTAGAACCCGCAATTATCCATCTAAAAAATTTGACAATATGGTGCTTGGTCCGGGTCAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_P18N2	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
E_P18N2_1	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
E_P18N2_3	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
E_P18N2_4	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
E_P18N2_2	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
Consensus	ACTATTCTACCAATAACAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTTCAAATTTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAAACTTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_P18N2	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P18N2_1	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P18N2_3	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P18N2_4	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P18N2_2	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_P18N2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
E_P18N2_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
E_P18N2_3	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
E_P18N2_4	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
E_P18N2_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAAATTGGGAATGTCCTGATTTTTTTCCTGTATCATTAAAAAATACAAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_P18N2	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCTTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
E_P18N2_1	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCTTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
E_P18N2_3	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCTTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
E_P18N2_4	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCTTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
E_P18N2_2	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCCTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
Consensus	TTAGATGCATCGTATCGCGGAAAAAATGTCAATATGTCCCTAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTCGATACCAATT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_P18N2	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_P18N2_1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_P18N2_3	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_P18N2_4	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_P18N2_2	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
Consensus	CTATCGATGGTTCGAAGGGATTGAGGCTTGACATATGGGAATTTCTATGCATCTAAATCATTCTATGACCTATGAAGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_P18N2	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
E_P18N2_1	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
E_P18N2_3	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
E_P18N2_4	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
E_P18N2_2	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
Consensus	TGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCTTAGTGGTAACAACTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATTCAA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_P18N2	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
E_P18N2_1	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
E_P18N2_3	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
E_P18N2_4	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
E_P18N2_2	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
Consensus	TTGAACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTTAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTGCTTCTCTTTTCAGTTTGAACAGGCCGACAAATTTGATCCTAAATGGG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
E_P18N2	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_P18N2_1	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_P18N2_3	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_P18N2_4	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_P18N2_2	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
Consensus	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTTCGACTATCCAGGTTGGGCTTGGACCATTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
E_P18N2	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_P18N2_1	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_P18N2_3	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_P18N2_4	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_P18N2_2	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
Consensus	TAGGCTCAAAAGAATTATAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
E_P18N2	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
E_P18N2_1	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
E_P18N2_3	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
E_P18N2_4	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
E_P18N2_2	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
Consensus	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGGTGATCCACGCTTAGCGATTCTAATAATATGCACATTTATTTGTCTTCAATA													
	1691	1700	1710	1720	1730	1740	1750	1758						
E_P18N2	ATGGATCTGAGACAAATCACAAATTTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTGGCTTCTCATATATATTTTGGACTTGCATCTTCAA													
E_P40N1_2	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTGGCTTCTCATATATATTTTGGACTTGCATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTGGCTTCTCATATATATTTTGGACTTGCATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA													
E_P40N1_2	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA													
Consensus	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_P40N1_2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCACTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	ACTATTCTACCAATAACAACCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
E_P40N1_2	ACTATTCTACCAATAACAACCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
Consensus	ACTATTCTACCAATAACAACCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTTATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	ATAACAACCCATTGATTGTACCTGACAATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAACCATAGAGGGAT													
E_P40N1_2	ATAACAACCCATTGATTGTACCTGACAATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAACCATAGAGGGAT													
Consensus	ATAACAACCCATTGATTGTACCTGACAATAGCATCAACAAACCAATTTTCGTGATCCACACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAACCATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGACCAAGCCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTGATTTTTCCCTGTATCATTAAAAATACTAATGGC													
E_P40N1_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGACCAAGCCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTGATTTTTCCCTGTATCATTAAAAATACTAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGACCAAGCCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTGATTTTTCCCTGTATCATTAAAAATACTAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	TTAGATGCATCGTATCGCGGAAAAAAAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACA													
E_P40N1_2	TTAGATGCATCGTATCGCGGAAAAAAAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACA													
Consensus	TTAGATGCATCGTATCGCGGAAAAAAAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	ATTCTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGA													
E_P40N1_2	ATTCTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGA													
Consensus	ATTCTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAGGAATCGAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	CGATGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATT													
E_P40N1_2	CGATGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATT													
Consensus	CGATGAATTAAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGATATGGCTCGACCCTAGTGGTAACAATTGATTCAATGGCCTATTGAAGAATTAGAACATTAGAAGCAAAAGATTATT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_P40N1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
E_P40N1_1	CAATTGAACAAACAGAAGTTGAGCAAGGGAGAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_P40N2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
E_P40N2_4	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
E_P40N2_2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
E_P40N2_1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
E_P40N2_3	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAARTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAAGGTGTTTGCTTCTCATATATTTTTTGGACTTGCATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_P40N2	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
E_P40N2_4	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
E_P40N2_2	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
E_P40N2_1	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
E_P40N2_3	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
Consensus	GTGCTATTAGTGTCAGAAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAAGACCTTAATGCACCAATGTATTATATGAGGTATATCATTTATTCTATCAATACACCCAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_P40N2	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_P40N2_4	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_P40N2_2	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_P40N2_1	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
E_P40N2_3	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
Consensus	AGGATCAGTATGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTAGAACCCGAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_P40N2	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
E_P40N2_4	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
E_P40N2_2	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
E_P40N2_1	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
E_P40N2_3	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
Consensus	ACTATTCTACCTAATAACAAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAAGATTATGCAATCCCGGCTAACTTGTCTGATCCATTCTTCGTAATGGATCAACCTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_P40N2	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P40N2_4	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P40N2_2	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P40N2_1	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P40N2_3	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACACCCGTTGATTGTTCTTGCAATAGCATCAACAAACCGAATTTCGTGATCCACACCCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTATAATAGGAAGTATGAGAAACATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_P40N2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
E_P40N2_4	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
E_P40N2_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
E_P40N2_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
E_P40N2_3	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCTTGATTTTTTCTCTGATCATTAAAAAATACTAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_P40N2	TTAGATGCATCGTATCGCGAAAAAATGTCAAACATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAA													
E_P40N2_4	TTAGATGCATCGTATCGCGAAAAAATGTCAAACATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGATTATT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_P54N1	-----													
E_P54N1_1	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
Consensus	ATGGAATTATTTATGAAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCCTCTTTATAGTTTTATCAACATTAAATGGGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA													
E_P54N1	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_P54N1_1	-----													
E_P54N1_1	GTGCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAA													
Consensus	GTGCTATTAGTGTCAAGAATGTTTCATAGAACTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTTATTCTATCAATACAATCCAA													
E_P54N1	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_P54N1_1	-----													
E_P54N1_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTGAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTGAGCA													
E_P54N1	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_P54N1_1	-----													
E_P54N1_1	ACTATTCTACCAATAACAAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
Consensus	ACTATTCTACCAATAACAAGCCCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAAGTTCAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA													
E_P54N1	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_P54N1_1	-----													
E_P54N1_1	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCAACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTACCTGACATAGCATCAACAAACCAATTTTCGTGATCCAACAACCGCATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGAGGGAT													
E_P54N1	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_P54N1_1	-----													
E_P54N1_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCCTGATTTTTTTCTGTATCATTAAAAAATACAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAAATGGGCCAAGGCCAACATCCACTTCATTTCATCTCCTCATACTGGAATTGGGAATGTCCTGATTTTTTTCTGTATCATTAAAAAATACAATGGC													
E_P54N1	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_P54N1_1	-----													
E_P54N1_1	TTAGATGCATCGTATCGCGGAAAAATGTCAAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
Consensus	TTAGATGCATCGTATCGCGGAAAAATGTCAAATATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGAGTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATAACAATT													
E_P54N1	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_P54N1_1	-----													
E_P54N1_1	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAAGAATCGAAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
Consensus	CTATCGATGGTTCGAAGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCATGAAGAATCGAAGAATTGTATGGGGTTGGACAATGAATCAGATGTTTTACCTGACGA													
E_P54N1	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_P54N1_1	-----													
E_P54N1_1	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCTAGTGGTAACAACCTGATTCAATGGCCTATTGAAGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
Consensus	TGAATTAAAGAAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCTAGTGGTAACAACCTGATTCAATGGCCTATTGAAGAATTAGAACATTAAAGAAGCAAAGATTATTCAA													
E_P54N1	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_P54N1_1	-----													
E_P54N1_1	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTCAAGTTTGACAAGGCCGAACAATTTGATCCTAATGGG													
Consensus	TTGAACAACAAGAGTTGAGCAAGGGAGAAATGTTTGAAGTTAAAGGAATCTCAGCATCACAGGCTGATATTGAAGTGTCTGTTCTCTTTTCAAGTTTGACAAGGCCGAACAATTTGATCCTAATGGG													
E_P54N1	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
E_P54N1_1	-----													
E_P54N1_1	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
Consensus	CCGACCTTTATGCCCAAGATGTTTGTGCCATAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGTTGGGCTTGTGACATTAGCTTCTAAAAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTT													
E_P54N1	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
E_P54N1_1	-----													
E_P54N1_1	TAAGGCTCAAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
Consensus	TAAGGCTCAAAAGAATTATAAGGTTCTCATGTGCTCAGATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTGCTGGATATGTAGATGTAGATTTAGTAGACACGAGAGAG													
E_P54N1	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
E_P54N1_1	-----													
E_P54N1_1	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGTATCCAACGTTAGCGATTACAAATATGCACATTTATTTGTCTTCAATA													
Consensus	TTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGTATCCAACGTTAGCGATTACAAATATGCACATTTATTTGTCTTCAATA													
E_P54N1	1691	1700	1710	1720	1730	1740	1750	1758						
E_P54N1_1	-----													
E_P54N1_1	ATGGATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	ATGGATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTATTAAAGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA														
E_P54N2_1	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTATTAAAGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA														
Consensus	ATGGAATTATTTATGAAAGCTCTTCTCTTTGGGGTTTAGAAATTTATTTATTTTGCTTCTTTATAGTTTTATCAACATTATTAAAGGTGTTTGCTTCTCATATATTTTTTTGGACTTGCAATCTTCAA														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAAAATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
E_P54N2_1	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAAAATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
Consensus	GTGCTATTAGTGTCAAGAAATGTTTCATAGAACTGGTTTTTCATTTTCACCTCCTAAAAATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAATCCAAA														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
E_P54N2_1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGCTTGGTCCGGGTCAGCA														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAGAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
E_P54N2_1	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAGAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
Consensus	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGGTTTCAGAAATTATGCAATCCCGGCTAACTTGTCTGATCCATTTCTTCGTAATGGATCAACCTA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	ATAACAACCCGTTGATTGTTCTTGACAAATAGCATCAACAACCCGAATTTTCGTGATCCAACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGGGGGAT														
E_P54N2_1	ATAACAACCCGTTGATTGTTCTTGACAAATAGCATCAACAACCCGAATTTTCGTGATCCAACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGGGGGAT														
Consensus	ATAACAACCCGTTGATTGTTCTTGACAAATAGCATCAACAACCCGAATTTTCGTGATCCAACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAATAGGAAGTATGAGAAACATAGGGGGAT														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAAAAATACTAATGGC														
E_P54N2_1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAAAAATACTAATGGC														
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGGCCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTATCATTAAAAAATACTAATGGC														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
E_P54N2_1	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
Consensus	TTAGATGCATCGTATCGCGAAAAAATGTCAACATGTCCTTAAGAATAGCCTTGATGTTAATAGGTTTGATTATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCTTGATACAATT														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	CTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTTTATGACCCTATGAGAATCGAGAATTGTATGGGGTTGGACAAATGAATCAGATGGTTTACCTGACGA														
E_P54N2_1	CTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTTTATGACCCTATGAGAATCGAGAATTGTATGGGGTTGGACAAATGAATCAGATGGTTTACCTGACGA														
Consensus	CTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTTTATGACCCTATGAGAATCGAGAATTGTATGGGGTTGGACAAATGAATCAGATGGTTTACCTGACGA														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
E_P54N2_1	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
Consensus	TGAATTAGAAGGATGGGCTGGAATTCAGCTATTCCGCGTAAGTATGGCTCGACCCTAGTGGTAACAATTGATTCATGGCCTATTGAAGAATTAGAACATTAGAAGGCAAAAGATTCAATTG														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACACAGGCCGAACAAATTTGATCCTAATGGGCCG														
E_P54N2_1	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACACAGGCCGAACAAATTTGATCCTAATGGGCCG														
Consensus	AACACACAGAGTTGAGCAAGGGAGAAATGTTTGAGTTGAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCGTTCTCTTTTTCAAGTTTGACACAGGCCGAACAAATTTGATCCTAATGGGCCG														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	ACCTTTATGCCAAGATGTTTGTCGTCATTAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACATTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
E_P54N2_1	ACCTTTATGCCAAGATGTTTGTCGTCATTAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACATTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
Consensus	ACCTTTATGCCAAGATGTTTGTCGTCATTAAAGGGTTCGACTATCCAAGGTGGGCTTGGACCATTGGGGCTTGCACATTAGCTTCTAAGAACTTGGAGAATACACACCTGTTTTCTTTCGAGTGTTTAA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAAGAAGTTA														
E_P54N2_1	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAAGAAGCTA														
Consensus	GGCTCAAAGAATTATAAGGTTCTCATGTGCTCAATGCTAGAAGATCTACCATGAGACAAATGAAGCAATGTACAGCCCTCATTTGCTGGATATGTAGATGTAGATTTAGTAGACATGAAGAAGTTA														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCAATGTTAGCGATTCAATAATGCACATTTATTTGTCTTCAATAATG														
E_P54N2_1	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCAATGTTAGCGATTCAATAATGCACATTTATTTGTCTTCAATAATG														
Consensus	TCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAAACATGCATAACATCGAGGGTGATCCAATGTTAGCGATTCAATAATGCACATTTATTTGTCTTCAATAATG														
	1691	1700	1710	1720	1730	1740	1750	1755							
E_P54N2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_P54N2_2	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
E_P54N2_1	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														
Consensus	GATCTGAGACAATCACAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
E_P18N1	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
E_P40N1	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
E_P18N2	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
E_P54N1	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
E_P40N2	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
E_P54N2	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
Consensus	ATGGAATTATTTATGAARAAGCTCTTCTCTTTGGGGTTTAGAATTTATTTATTTTGCCTCTTTATAGTTTATCAAAACATTAAATGGGGTGTGGTCTCTCATATATATTTTTTGGAGCTTGCATCTTCAA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
E_P18N1	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P40N1	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P18N2	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAATATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P54N1	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P40N2	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
E_P54N2	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
Consensus	GTGCTATTAGTGTCAGGAATGTTCATAGAAGCTGGTTTTTCATTTTCAACCTCCTAAACATTGGATTAAATGACCCTAATGCACCAATGTATTATAATGGAGTATATCATTATTCTATCAATACAAATCCAA													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
E_P18N1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
E_P40N1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
E_P18N2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
E_P54N1	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
E_P40N2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
E_P54N2	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
Consensus	AGGATCAGTATGGGGCAATATTGTTTGGGCTCATTCAGTCTCAAAGACTTGATAAATTGGATCCATTTAGAACCCGCAATTTATCCATCTAAAAAATTTGACAAATATGGTGTCTGGTCCGGGTCAGCA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
E_P18N1	ACTATTCTACCAATAACAACCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
E_P40N1	ACTATTCTACCAATAACAACCCATTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
E_P18N2	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
E_P54N1	ACTATTCTACCAATAACAAGCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
E_P40N2	ACTATTCTACCTAATAACAACCTGTTATCTTATATACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
E_P54N2	ACTATTCTACCTAATAACAACCTGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
Consensus	ACTATTCTACCAATAACAACCCGTTATCTTATACACCGGAGTAGTAGATTCCCATGATTCTCAGTTCAAATATGCAATCCCGGCTAAGTTGTCTGATCCATTTCTTCGTAATGGATCAACCTTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
E_P18N1	ATAACAACCCATTGATTGTACCTGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
E_P40N1	ATAACAACCCATTGATTGTACCTGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
E_P18N2	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
E_P54N1	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
E_P40N2	ATAACAACCCGTTGATTGTCTCGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
E_P54N2	ATAACAACCCGTTGATTGTCTCGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
Consensus	ATAACAACCCGTTGATTGTACCTGACAAATAGCATCAACAACCAATATTCGTGATCCACAACCCGATGGATGGGCCAAGATGGGCTTTGGAGAATTGTAAATAGGAAGTATGAGAAACCATAGAGGGAT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
E_P18N1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_P40N1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_P18N2	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_P54N1	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_P40N2	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
E_P54N2	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
Consensus	GGCATTATTGTATAGAAGTAGAGACTTCATTAATGGACCAAGGCCAACATCCACTTCATTCATCTCCTCATACTGGAATTTGGGAATGTCTTGATTTTTTCCCTGTATCATTAAAAAATACTAATGGC													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
E_P18N1	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
E_P40N1	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
E_P18N2	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
E_P54N1	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
E_P40N2	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
E_P54N2	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
Consensus	TTAGATGCATCGTATCGCGGAAAAAATAATGTCAACATGTCTTAAAGATAGCCTTGATGTTAATAGGTTTGAATATTACACTATTGGTATGTATGACACCAAAAAAGATAGGTACATTCCCTGATACCA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
E_P18N1	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
E_P40N1	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
E_P18N2	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
E_P54N1	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
E_P40N2	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
E_P54N2	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
Consensus	ATTCTATCGATGGTTCGAGGGGATTGAGGCTTGACTATGGGAATTTCTATGCATCTAATCATTCTATGACCCCTATGAGGAATCGAGGAATTTGATGGGGTTGGACAAATGAATCAGATGTTTTACCTGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
E_P18N1	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATTATT													
E_P40N1	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATTATT													
E_P18N2	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATTATT													
E_P54N1	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATTATT													
E_P40N2	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACTTCTAGTGTTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATT---													
E_P54N2	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATT---													
Consensus	CGATGAATTAAGAAAGGATGGGCTGGAAATTCAGCTATTCCGCGTAAAGTATGGCTCGACCCCTAGTGGTAAACAATTGATTCAATGGCCTATTGAGGAATTAGAAACATTAAAGAAAGCAAAGATTat													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
E_P18N1	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
E_P40N1	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
E_P18N2	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
E_P54N1	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
E_P40N2	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
E_P54N2	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
Consensus	CAATTGAACACACAGAGTTGAGCAGGGGAGAAATGTTTGAGGTTAAGGGAATCTCAGCATCACAGGCTGATATTGAGTGTCTCTCTTTTCAGGTTTAACAGGGCCGACAAATTTGATCCTAAT													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
E_P18N1	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAAAACCTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
E_P40N1	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAAAACCTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
E_P18N2	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAAAACCTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
E_P54N1	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAAAACCTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
E_P40N2	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAGAACTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
E_P54N2	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAGAACTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
Consensus	GGGCCGACCTTTATGCCAAGATGTTTGTCATTAAGGGTTTCGACTATCCAAGGTGGGCTTGGACCATTTGGGCTTGGACATTAGCTTCTAAAAACCTTGGAGAGATACACACCTGTTTTCTTTCGAGT													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
E_P18N1	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
E_P40N1	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
E_P18N2	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
E_P54N1	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
E_P40N2	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
E_P54N2	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
Consensus	GTTTAAGGCTCAAAAGAAATTAAGGTTCTCATGTGCTCAGATGCTAGAGATCTACCATGAGACAAATGAGCAATGTACAGCCCTCATTTCGCTGGATATGTAGATGTAGATTAGTAGACACGAG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
E_P18N1	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
E_P40N1	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
E_P18N2	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
E_P54N1	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
E_P40N2	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
E_P54N2	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
Consensus	AAGTTATCTCTTAGGAGTTTGATTGATAACTCAGTAGTGGAGAGTTTTGGTGCTGGTGGAAAACATGCATACATCGAGGGTGATCCACGTTAGCGATTCAATAARTGCACATTTATTTGTCTTCA													
	1691	1700	1710	1720	1730	1740	1750	1761						
E_P18N1	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_P40N1	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_P18N2	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_P54N1	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_P40N2	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
E_P54N2	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													
Consensus	ATAATGGATCTGAGACATCACAAATTGAGACTCTTAATGCTTGGAGCATGGATGTACCTAAGATGCACTAA													

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
		+	+	+	+	+	+	+	+	+	+	+	+	+
F_TN1	MDYSSNSRWALPVILVCFVYLLSNMVVFASHKVFIHLQSQNAVNVHTVHRTGYHFQPEKHWINDPNAPMYFNGVYHLFYQYNPNGSVMGNIYWAHSVSKDLINWINLEPAIYPSKPFQDQFGTWSGSATI													
F_TN1_1	MDYSSNSRWALPVILVCFVYLLSNMVVFASHKVFIHLQSQNAVNVHTVHRTGYHFQPEKHWINDPNAPMYFNGVYHLFYQYNPNGSVMGNIYWAHSVSKDLINWINLEPAIYPSKPFQDQFGTWSGSATI													
F_TN2	MDYSSNSRWALPVILVCFFIYLLSNMVVFASHKVFIHLQSQNAVNVQTVHRTGYHFQPEKHWINDPNAPMYFNGVYHLFYQYNPNGSVMGNIYWAHSVSKDLINWINLEPAIYPSKPFQDQFGTWSGSATI													
Consensus	MDYSSNSRWALPVILVCF !YLLSNMVVFASHKVFIHLQSQNAVNVhTVHRTGYHFQPEKHWINDPNAPMYFNGVYHLFYQYNPNGSVMGNIYWAHSVSKDLINWINLEPAIYPSKPFQDQFGTWSGSATI													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
		+	+	+	+	+	+	+	+	+	+	+	+	+
F_TN1	LPGNKPVILYTGIVDANQTQVQNYAVPANISDPYLREWIKPDNNPLIYADASINKTKFRDPTTAWMGKDGHWRIYMGSLRKHSRGLAIMYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPYALKGTNGQ													
F_TN1_1	LPGNKPVILYTGIVDANQTQVQNYAVPANISDPYLREWIKPDNNPLIYADASINKTKFRDPTTAWMGKDGHWRIYMGSLRKHSRGLAIMYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPYALKGTNGQ													
F_TN2	LPGNKPVILYTGIVDANQTQVQNYAVPANLSDPHLREWIKPDNNPLIYADASINKTKFRDPTTAWMGKDGHWRIYMGSLRKHSRGLAIMYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPYALKGTNGI													
Consensus	LPGNKPVILYTGIVDANQTQVQNYAVPANiSDPyLREWIKPDNNPLIYADASINKTKFRDPTTAWMGKDGHWRIYMGSLRKHSRGLAIMYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPYALKGTNGq													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
		+	+	+	+	+	+	+	+	+	+	+	+	+
F_TN1	DQYGEEHKYVLKNSMDLTRFEYITLGKYDTKKDRYIPDVGSIDSNKGLRFDYGNFYASKSFYDPSKNRRVINGWSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLIQWPVEELETLRQKVQLSNK													
F_TN1_1	DQYGEEHKYVLKNSMDLTRFEYITLGKYDTKKDRYIPDVGSIDSNKGLRFDYGNFYASKSFYDPSKNRRVINGWSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLIQWPVEELETLRQKVQLSNK													
F_TN2	DQYGEEYKNVLKNGMDLTRFEYITLGKYDTKKDRYVPDVGSIDSNKGLRFDYGNFYASKTFYDTSKNRRVINGWSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLIQWPVEELETLRQKVQLSNK													
Consensus	DQYGEEhKyVLKNsMDLTRFEYITLGKYDTKKDRY!PDVGSIDSNKGLRFDYGNFYASKsFYDpSKNRRVINGWSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLIQWPVEELETLRQKVQLSNK													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
		+	+	+	+	+	+	+	+	+	+	+	+	+
F_TN1	KLNGEKVEYTGITPAQADVEYTFASFSLDKAESFDPsWTDHYAQDYCGLKGADYQGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKVLLCSDAKRSTLKFNETHYkVSFAGFYDVLDADKKLSLRS													
F_TN1_1	KLNGEKVEYTGITPAQADVEYTFASFSLDKAESFDPsWTDHYAQDYCGLKGADYQGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKVLLCSDAKRSTLKFNETHYRVSFAGFYDVLDADKKLSLRS													
F_TN2	KLNGEKVEYTGITPAQADVEYTFASFSLDKAELFDSSWTDHYAQDYCGLKGADYQGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKVLLCSDAKRSTLKFNETHYkVSFAGFYDVLDADKKLSLRS													
Consensus	KLNGEKVEYTGITPAQADVEYTFASFSLDKAESFDpsWTDHYAQDYCGLKGADYQGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKVLLCSDAKRSTLKFNETHYkVSFAGFYDVLDADKKLSLRS													
	521	530	540	550	560	570	581							
		+	+	+	+	+	+							
F_TN1	LIDNSVIESFGAGGKTCITSRVYPTLAINDKAHLFAFNNGTEPITIETLDAWSHGKAKIQY													
F_TN1_1	LIDNSVIESFGAGGKTCITSRVYPTLAINDKAHLFAFNNGTEPITIETLDAWSHGKAKIQY													
F_TN2	LIDNSVIESFGAGGKTCITSRVYPTLAINEKAHLFAFNNGTEPITIETLDAWSHGKAKIQY													
Consensus	LIDNSVIESFGAGGKTCITSRVYPTLAIN#KAHLFAFNNGTEPITIETLDAWSHGKAKIQY													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
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F_SN1	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTTCATTTCGAGTCTCAAAATGCTG														
F_SN1_1	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTTCATTTCGAACTCTCAAAATGCTG														
F_SN1_2	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTTCATTTCGAACTCTCAAAATGCTG														
Consensus	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTTCATTTCGAaTCTCAAAATGCTG														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	TAAATGTTCAAACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC														
F_SN1_1	TAAATGTTCAAACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC														
F_SN1_2	TAAATGTTCAAACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC														
Consensus	TAAATGTTCAAACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT														
F_SN1_1	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT														
F_SN1_2	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT														
Consensus	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	CTACCTGGTAAACAGCCTGTTATCTTGACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAAACTACGCGATTCAGCTAAGCTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
F_SN1_1	CTACCTGGTAAACAGCCTGTTATCTTGACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAAACTACGCGAATTCAGCTAAGCTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
F_SN1_2	CTACCTGGTAAACAGCCTGTTATCTTGACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAAACTACGCGATTCAGCTAAGCTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
Consensus	CTACCTGGTAAACAGCCTGTTATCTTGACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAAACTACGCGaTTCAGCTAAGCTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
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F_SN1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAAGACCAAAATTCGTGATCCACACACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC														
F_SN1_1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAAGACCAAAATTCGTGATCCACACACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC														
F_SN1_2	ATCCATTGATTGTAGCTGATGCTAGCATCAACAAGACCAAAATTCGTGATCCACACACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC														
Consensus	ATCCATTGATTGTAGCTGATGCTAGCATCAACAAGACCAAAATTCGTGATCCACACACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
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F_SN1	TATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCTGATTTTTTCCCTGTTGCATTGAAGGGAAGTAATGGGATA														
F_SN1_1	TATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCTGATTTTTTCCCTGTTGCATTGAAGGGAAGTAATGGGATA														
F_SN1_2	TATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCTGATTTTTTCCCTGTTGCATTGAAGGGAAGTAATGGGATA														
Consensus	TATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCTGATTTTTTCCCTGTTGCATTGAAGGGAAGTAATGGGATA														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	GATCAATATGGTGAAGAAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAGGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
F_SN1_1	GATCAATATGGTGAAGAAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAGGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
F_SN1_2	GATCAATATGGTGAAGAAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAGGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
Consensus	GATCAATATGGTGAAGAAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAGGATAGGTATGTTCCAGATGTTGGTTCTATTGATa														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC														
F_SN1_1	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC														
F_SN1_2	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC														
Consensus	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	GAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACAATTGGTTCAAGTGGCCTGTTGAAGAATTAGAACCCTAAGAACCCAAAGGTTCAATTGAGTAACAAA														
F_SN1_1	GAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCAAAGGTTCAATTGAGTAACAAA														
F_SN1_2	GAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCAAAGGTTCAATTGAGTAACAAA														
Consensus	GAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGaTCCAGTGGAACAATTGGTTCAaTGGCCTGTTGAAGAATTAGAACCCTAAGAACCCAAAGGTTCAATTGAGTAACAAA														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	AAATTGAACAATGGTGAAAAGGTTGAAGTTACAGGAATCACACCTGCACAGGCGAGATGTTGAAGTGACATTCTCTTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
F_SN1_1	AAATTGAACAATGGTGAAAAGGTTGAAGTTACAGGAATCACACCTGCACAGGCGGATGTTGAAGTGATATTTCTTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
F_SN1_2	AAATTGAACAATGGTGAAAAGGTTGAAGTTACAGGAATCACACCTGCACAGGCGGATGTTGAAGTGATATTTCTTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
Consensus	AAATTGAACAATGGTGAAAAGGTTGAAGTTACAGGAATCACACCTGCACAGGCGGATGTTGAAGTGATATTTCTTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	CACAGATGTTTGTGGACTCAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTTCTTCCGAGTTTCAAGCACAAACA														
F_SN1_1	CACAGATGTTTGTGGACTTAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTTCTTCCGAGTTTCAAGCACAAACA														
F_SN1_2	CACAGATGTTTGTGGACTTAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTTCTTCCGAGTTTCAAGCACAAACA														
Consensus	CACAGATGTTTGTGGACTTAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTTCTTCCGAGTTTCAAGCACAAACA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCAACTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCAGACAAGAATTGTCACTCAGAAGC														
F_SN1_1	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCAACTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAAGC														
F_SN1_2	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCAACTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAAGC														
Consensus	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCAACTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAAGC														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACACGGAAGCTGAGCCAA														
F_SN1_1	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCAATTCACACGGAAGCTGAGCCAA														
F_SN1_2	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCAATTCACACGGAAGCTGAGCCAA														
Consensus	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCaTTCACACGGAAGCTGAGCCAA														
	1691	1700	1710	1720	1730	1740	1746								
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
F_SN1	TCACAATTGAGACTTTGGATGcATGGAGTATGGGCAAGCTAAGATACAATATTGA														
F_SN1_1	TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTAAGATACAATATTGA														
F_SN1_2	TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTAAGATACAATATTGA														
Consensus	TCACAATTGAGACTTTGGATGcATGGAGTATGGGCAAGCTAAGATACAATATTGA														

1 10 20 30 40 50 60 70 80 90 100 110 120 130
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTTATTCACTTGCAATCTCAAATGCTG
F_SN2_1 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTTATTCACTTGCAATCTCAAATGCTG
F_SN2_3 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTTATTCACTTGCAATCTCAAATGCTG
F_SN2_2 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTTATTCACTTGCAATCTCAAATGCTG
Consensus ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTTATTCACTTGCAATCTCAAATGCTG

131 140 150 160 170 180 190 200 210 220 230 240 250 260
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 TAAATGTTCCACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAAATTTACCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN2_1 TAAATGTTCCACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAAATTTACCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN2_3 TAAATGTTCCACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAAATTTACCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN2_2 TAAATGTTCCACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAAATTTACCATCTATTCTACCAATACAACCCAAACGGGTC
Consensus TAAATGTTCCACTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAAATTTACCATCTATTCTACCAATACAACCCAAACGGGTC

261 270 280 290 300 310 320 330 340 350 360 370 380 390
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F_SN2 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAAACCATTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT
F_SN2_1 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAAACCATTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT
F_SN2_3 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAAACCATTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT
F_SN2_2 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAAACCATTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT
Consensus AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAAACCATTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT

391 400 410 420 430 440 450 460 470 480 490 500 510 520
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F_SN2 CTACCTGGTAAACAGCCTGTTATCTTGTACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAATTTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAAGCCTGATAACA
F_SN2_1 CTACCTGGTAAACAGCCTGTTATCTTGTACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAATTTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAAGCCTGATAACA
F_SN2_3 CTACCTGGTAAACAGCCTGTTATCTTGTACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAATTTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAAGCCTGATAACA
F_SN2_2 CTACCTGGTAAACAGCCTGTTATCTTGTACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAATTTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAAGCCTGATAACA
Consensus CTACCTGGTAAACAGCCTGTTATCTTGTACACTGGAATAGTAGATGCCAACCAAACTCAAGTTCAAATTTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAAGCCTGATAACA

521 530 540 550 560 570 580 590 600 610 620 630 640 650
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F_SN2 ATCCATTGATTGTAGCTGATGATAGTATCAACAAACCAAAATTTCTGATCCAACTACTGCTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC
F_SN2_1 ATCCATTGATTGTAGCTGATGATAGTATCAACAAACCAAAATTTCTGATCCAACTACTGCTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC
F_SN2_3 ATCCATTGATTGTAGCTGATGATAGTATCAACAAACCAAAATTTCTGATCCAACTACTGCTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC
F_SN2_2 ATCCATTGATTGTAGCTGATGATAGTATCAACAAACCAAAATTTCTGATCCAACTACTGCTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC
Consensus ATCCATTGATTGTAGCTGATGATAGTATCAACAAACCAAAATTTCTGATCCAACTACTGCTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC

651 660 670 680 690 700 710 720 730 740 750 760 770 780
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F_SN2 AATAATGTATAGAGCAAGATTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTGATTTTTTCTGTTGCATTGAAGGGAACCTAATGGGATA
F_SN2_1 AATAATGTATAGAGCAAGATTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTGATTTTTTCTGTTGCATTGAAGGGAACCTAATGGGATA
F_SN2_3 AATAATGTATAGAGCAAGATTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTGATTTTTTCTGTTGCATTGAAGGGAACCTAATGGGATA
F_SN2_2 AATAATGTATAGAGCAAGATTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTGATTTTTTCTGTTGCATTGAAGGGAACCTAATGGGATA
Consensus AATAATGTATAGAGCAAGATTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTGATTTTTTCTGTTGCATTGAAGGGAACCTAATGGGATA

781 790 800 810 820 830 840 850 860 870 880 890 900 910
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F_SN2 GATCAATATGGTGAAGAAATATAAATATGTGCTTAAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAAGTATGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTCTATTGATA
F_SN2_1 GATCAATATGGTGAAGAAATATAAATATGTGCTTAAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAAGTATGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTCTATTGATA
F_SN2_3 GATCAATATGGTGAAGAAATATAAATATGTGCTTAAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAAGTATGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTCTATTGATA
F_SN2_2 GATCAATATGGTGAAGAAATATAAATATGTGCTTAAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAAGTATGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTCTATTGATA
Consensus GATCAATATGGTGAAGAAATATAAATATGTGCTTAAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAAGTATGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTCTATTGATA

911 920 930 940 950 960 970 980 990 1000 1010 1020 1030 1040
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F_SN2 GTTGGAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGAGCTTTCTATGATACTAGCAAAAACCGAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTTGAGAGTGATAATGC
F_SN2_1 GTTGGAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGAGCTTTCTATGATACTAGCAAAAACCGAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTTGAGAGTGATAATGC
F_SN2_3 GTTGGAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGAGCTTTCTATGATACTAGCAAAAACCGAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTTGAGAGTGATAATGC
F_SN2_2 GTTGGAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGAGCTTTCTATGATACTAGCAAAAACCGAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTTGAGAGTGATAATGC
Consensus GTTGGAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGAGCTTTCTATGATACTAGCAAAAACCGAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTTGAGAGTGATAATGC

1041 1050 1060 1070 1080 1090 1100 1110 1120 1130 1140 1150 1160 1170
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGGTTCAAGTGGCTGTTGAAGAAATAGAAACCTAAGAACCCAAAGGTTCAATTGAGTAACAAA
F_SN2_1 GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGGTTCAAGTGGCTGTTGAAGAAATAGAAACCTAAGAACCCAAAGGTTCAATTGAGTAACAAA
F_SN2_3 GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGGTTCAAGTGGCTGTTGAAGAAATAGAAACCTAAGAACCCAAAGGTTCAATTGAGTAACAAA
F_SN2_2 GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGGTTCAAGTGGCTGTTGAAGAAATAGAAACCTAAGAACCCAAAGGTTCAATTGAGTAACAAA
Consensus GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGGTTCAAGTGGCTGTTGAAGAAATAGAAACCTAAGAACCCAAAGGTTCAATTGAGTAACAAA

1171 1180 1190 1200 1210 1220 1230 1240 1250 1260 1270 1280 1290 1300
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 AAATTGAACAAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG
F_SN2_1 AAATTGAACAAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG
F_SN2_3 AAATTGAACAAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG
F_SN2_2 AAATTGAACAAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG
Consensus AAATTGAACAAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG

1301 1310 1320 1330 1340 1350 1360 1370 1380 1390 1400 1410 1420 1430
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACAC
F_SN2_1 CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACAC
F_SN2_3 CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACAC
F_SN2_2 CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACAC
Consensus CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACAC

1431 1440 1450 1460 1470 1480 1490 1500 1510 1520 1530 1540 1550 1560
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 AAATTACAGGTTCTCTTGTTGTTCTGACGCTAAAGGTCACCTCTTAGGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN2_1 AAATTACAGGTTCTCTTGTTGTTCTGACGCTAAAGGTCACCTCTTAGGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN2_3 AAATTACAGGTTCTCTTGTTGTTCTGACGCTAAAGGTCACCTCTTAGGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN2_2 AAATTACAGGTTCTCTTGTTGTTCTGACGCTAAAGGTCACCTCTTAGGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
Consensus AAATTACAGGTTCTCTTGTTGTTCTGACGCTAAAGGTCACCTCTTAGGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC

1561 1570 1580 1590 1600 1610 1620 1630 1640 1650 1660 1670 1680 1690
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
F_SN2 TTGATTGATAATTCAGTTATAGAAGTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAGGACACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN2_1 TTGATTGATAATTCAGTTATAGAAGTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAGGACACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN2_3 TTGATTGATAATTCAGTTATAGAAGTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAGGACACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN2_2 TTGATTGATAATTCAGTTATAGAAGTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAGGACACATTTATTTGCGTTCAACACGGAACTGAGCCAA
Consensus TTGATTGATAATTCAGTTATAGAAGTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAGGACACATTTATTTGCGTTCAACACGGAACTGAGCCAA

1691 1700 1710 1720 1730 1740 1746
|-----|-----|-----|-----|-----|-----|-----|
F_SN2 TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA
F_SN2_1 TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA
F_SN2_3 TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA
F_SN2_2 TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA
Consensus TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_SN3	-----													
F_SN3_1	ATGGATTATTCATCTAATTCTCGTGGGCTTTGCCAGTTATCTTAGTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCATCTCAAATGCTG													
F_SN3_2	ATGGATTATTCATCTAATTCTCGTAGGGCTTTGCCAGTTATCTTAGTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCATCTCAAATGCTG													
F_SN3_3	ATGGATTATTCATCTAATTCTCGTGGGCTTTGCCAGTTATCTTAGTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCATCTCAAATGCTG													
Consensus	ATGGATTATTCATCTAATTCTCGTGGGCTTTGCCAGTTATCTTAGTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCATCTCAAATGCTG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_SN3	-----													
F_SN3_1	TAAATGTTCAAACGTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTTTACCATCTATTCTACCAATACACCCAAACGGGTC													
F_SN3_2	TAAATGTTCAAACGTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTTTACCATCTATTCTACCAATACACCCAAACGGGTC													
F_SN3_3	TAAATGTTCAAACGTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTTTACCATCTATTCTACCAATACACCCAAACGGGTC													
Consensus	TAAATGTTCAAACGTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTTTACCATCTATTCTACCAATACACCCAAACGGGTC													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_SN3	-----													
F_SN3_1	AGTATGGGGCAATATTGTTTGGGCTCATTACGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTCGGTACGTGGTCTGGGCTGCAACTATT													
F_SN3_2	AGTATGGGGCAATATTGTTTGGGCTCATTACGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTCGGTACGTGGTCTGGGCTGCAACTATT													
F_SN3_3	AGTATGGGGCAATATTGTTTGGGCTCATTACGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTCGGTACGTGGTCTGGGCTGCAACTATT													
Consensus	AGTATGGGGCAATATTGTTTGGGCTCATTACGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTCGGTACGTGGTCTGGGCTGCAACTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_SN3	-----													
F_SN3_1	CTACCTGGTAACAGCCTGTTATTTGTACACTGGAAATAGTAGATGCCAACCAAACTCAGTTCAAACCTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAGCCTGATAATA													
F_SN3_2	CTACCTGGTAACAGCCTGTTATTTGTACACTGGAAATAGTAGATGCCAACCAAACTCAGTTCAAACCTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAGCCTGATAATA													
F_SN3_3	CTACCTGGTAACAGCCTGTTATTTGTACACTGGAAATAGTAGATGCCAACCAAACTCAGTTCAAACCTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAGCCTGATAATA													
Consensus	CTACCTGGTAACAGCCTGTTATTTGTACACTGGAAATAGTAGATGCCAACCAAACTCAGTTCAAACCTACGCGATTCCAGCTAACGTATCTGATCCTTATCTCCGTGAATGGATCAGCCTGATAATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_SN3	-----													
F_SN3_1	ATCCATTGATTGTAGCTGATGATAGTATCACAAGACCAAAATTTTCGTGATCCACAACCTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
F_SN3_2	ATCCATTGATTGTAGCTGATGATAGTATCACAAGACCAAAATTTTCGTGATCCACAACCTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
F_SN3_3	ATCCATTGATTGTAGCTGATGATAGTATCACAAGACCAAAATTTTCGTGATCCACAACCTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
Consensus	ATCCATTGATTGTAGCTGATGATAGTATCACAAGACCAAAATTTTCGTGATCCACAACCTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_SN3	-----													
F_SN3_1	AATAATGTATAGAGCAAGATTTTATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTTGATTTTTTCTGTTGCATTGAAGGGAACATATGGGATA													
F_SN3_2	AATAATGTATAGAGCAAGATTTTATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTTGATTTTTTCTGTTGCATTGAAGGGAACATATGGGATA													
F_SN3_3	AATAATGTATAGAGCAAGATTTTATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTTGATTTTTTCTGTTGCATTGAAGGGAACATATGGGATA													
Consensus	AATAATGTATAGAGCAAGATTTTATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAATGGTACTGGAATTTGGGAATGTCTTGATTTTTTCTGTTGCATTGAAGGGAACATATGGGATA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_SN3	-----													
F_SN3_1	GATCAATATGGTGAGGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCATTGATA													
F_SN3_2	GATCAATATGGTGAGGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCATTGATA													
F_SN3_3	GATCAATATGGTGAGGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCATTGATA													
Consensus	GATCAATATGGTGAGGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCATTGATA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_SN3	-----													
F_SN3_1	GTTGGAGGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACCTTCTATGATCTAGCAAAAACCGAGGGGTATTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGATGATAATGC													
F_SN3_2	GTTGGAGGGGATTGAGATTCGATTATGGTAATTTCTATGATCTAGCAAAAACCGAGGGGTATTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGATGATAATGC													
F_SN3_3	GTTGGAGGGGATTGAGATTCGATTATGGTAATTTCTATGATCTAGCAAAAACCGAGGGGTATTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGATGATAATGC													
Consensus	GTTGGAGGGGATTGAGATTCGATTATGGTAATTTCTATGATCTAGCAAAAACCGAGGGGTATTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGATGATAATGC													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_SN3	-----													
F_SN3_1	GAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGAACCAATTGATTCAATGGCTGTTGAGGAATTAGAACCTTAGAACCCAAAGGTTCAATTG													

1 10 20 30 40 50 60 70 80 90 100 110 120 130
|-----|
F_SN4 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTGTAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCATAAAGTTTTATTCACTTGCAATCTCAAATGCTG
F_SN4_2 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTGTAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCATAAAGTTTTATTCACTTGCAATCTCAAATGCTG
F_SN4_3 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTGTAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCATAAAGTTTTATTCACTTGCAATCTCAAATGCTG
F_SN4_1 ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTGTAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCATAAAGTTTTATTCACTTGCAATCTCAAATGCTG
Consensus ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTGTAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCATAAAGTTTTATTCACTTGCAATCTCAAATGCTG

131 140 150 160 170 180 190 200 210 220 230 240 250 260
|-----|
F_SN4 TAAATGTTCCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN4_2 TAAATGTTCCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN4_3 TAAATGTTCCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCCAAACGGGTC
F_SN4_1 TAAATGTTCCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCCAAACGGGTC
Consensus TAAATGTTCCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTTGGATCAATGATCCCARTGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCCAAACGGGTC

261 270 280 290 300 310 320 330 340 350 360 370 380 390
|-----|
F_SN4 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACC CGCAATTTACCCATCTAAACCATTTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT
F_SN4_2 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACC CGCAATTTACCCATCTAAACCATTTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT
F_SN4_3 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACC CGCAATTTACCCATCTAAACCATTTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT
F_SN4_1 AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACC CGCAATTTACCCATCTAAACCATTTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT
Consensus AGTATGGGGCAACATTGTTTGGGCTCATTAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACC CGCAATTTACCCATCTAAACCATTTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT

391 400 410 420 430 440 450 460 470 480 490 500 510 520
|-----|
F_SN4 CTACCTGGTAAACAACCTGTTATCTTGTACACTGGAATAGTGGATGCTAACCAAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGGCTGATACCA
F_SN4_2 CTACCTGGTAAACAACCTGTTATCTTGTACACTGGAATAGTGGATGCTAACCAAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGGCTGATACCA
F_SN4_3 CTACCTGGTAAACAACCTGTTATCTTGTACACTGGAATAGTGGATGCTAACCAAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGGCTGATACCA
F_SN4_1 CTACCTGGTAAACAACCTGTTATCTTGTACACTGGAATAGTGGATGCTAACCAAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGGCTGATACCA
Consensus CTACCTGGTAAACAACCTGTTATCTTGTACACTGGAATAGTGGATGCTAACCAAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGGCTGATACCA

521 530 540 550 560 570 580 590 600 610 620 630 640 650
|-----|
F_SN4 ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTTCTGATCCACACACCGCTTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC
F_SN4_2 ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTTCTGATCCACACACCGCTTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC
F_SN4_3 ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTTCTGATCCACACACCGCTTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC
F_SN4_1 ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTTCTGATCCACACACCGCTTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC
Consensus ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTTCTGATCCACACACCGCTTGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC

651 660 670 680 690 700 710 720 730 740 750 760 770 780
|-----|
F_SN4 TATAATGTATAGAGCAAGACTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGTTGCATTGAAGGGAACCTAATGGGCAT
F_SN4_2 TATAATGTATAGAGCAAGACTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGTTGCATTGAAGGGAACCTAATGGGCAT
F_SN4_3 TATAATGTATAGAGCAAGACTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGTTGCATTGAAGGGAACCTAATGGGCAT
F_SN4_1 TATAATGTATAGAGCAAGACTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGTTGCATTGAAGGGAACCTAATGGGCAT
Consensus TATAATGTATAGAGCAAGACTTCATGAATGGGTTAAGGCTAAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGTTGCATTGAAGGGAACCTAATGGGCAT

781 790 800 810 820 830 840 850 860 870 880 890 900 910
|-----|
F_SN4 GATCAATATGGTGAAGACACAAATATGTGCTTAAAGATAGTATGGATCTTACTCGATTGAGTACTATACTCTTGGTAAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTCTATTGATA
F_SN4_2 GATCAATATGGTGAAGACACAAATATGTGCTTAAAGATAGTATGGATCTTACTCGATTGAGTACTATACTCTTGGTAAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTCTATTGATA
F_SN4_3 GATCAATATGGTGAAGACACAAATATGTGCTTAAAGATAGTATGGATCTTACTCGATTGAGTACTATACTCTTGGTAAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTCTATTGATA
F_SN4_1 GATCAATATGGTGAAGACACAAATATGTGCTTAAAGATAGTATGGATCTTACTCGATTGAGTACTATACTCTTGGTAAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTCTATTGATA
Consensus GATCAATATGGTGAAGACACAAATATGTGCTTAAAGATAGTATGGATCTTACTCGATTGAGTACTATACTCTTGGTAAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTCTATTGATA

911 920 930 940 950 960 970 980 990 1000 1010 1020 1030 1040
|-----|
F_SN4 GTTGGAGGGGATTGAGATTGCACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAACGC
F_SN4_2 GTTGGAGGGGATTGAGATTGCACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAACGC
F_SN4_3 GTTGGAGGGGATTGAGATTGCACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAACGC
F_SN4_1 GTTGGAGGGGATTGAGATTGCACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAACGC
Consensus GTTGGAGGGGATTGAGATTGCACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAACGC

1041 1050 1060 1070 1080 1090 1100 1110 1120 1130 1140 1150 1160 1170
|-----|
F_SN4 TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACCAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCTAAGGTTCAATTGAGCAACAG
F_SN4_2 TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACCAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCTAAGGTTCAATTGAGCAACAG
F_SN4_3 TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACCAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCTAAGGTTCAATTGAGCAACAG
F_SN4_1 TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACCAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCTAAGGTTCAATTGAGCAACAG
Consensus TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACCAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAAGAACCCTAAGGTTCAATTGAGCAACAG

1171 1180 1190 1200 1210 1220 1230 1240 1250 1260 1270 1280 1290 1300
|-----|
F_SN4 AAGTTGAACAAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAGGCGAGAGTCATTTGATCCTAGTTGGACTGATATGTATG
F_SN4_2 AAGTTGAACAAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAGGCGAGAGTCATTTGATCCTAGTTGGACTGATATGTATG
F_SN4_3 AAGTTGAACAAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAGGCGAGAGTCATTTGATCCTAGTTGGACTGATATGTATG
F_SN4_1 AAGTTGAACAAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAGGCGAGAGTCATTTGATCCTAGTTGGACTGATATGTATG
Consensus AAGTTGAACAAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTTGCAGTTTGGATAGGCGAGAGTCATTTGATCCTAGTTGGACTGATATGTATG

1301 1310 1320 1330 1340 1350 1360 1370 1380 1390 1400 1410 1420 1430
|-----|
F_SN4 CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACCA
F_SN4_2 CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACCA
F_SN4_3 CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACCA
F_SN4_1 CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACCA
Consensus CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACCTTAGAAGAAAACACACCTGTTTCTTCCGAGTTTTCAAAGCACACCA

1431 1440 1450 1460 1470 1480 1490 1500 1510 1520 1530 1540 1550 1560
|-----|
F_SN4 AAATTACAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN4_2 AAATTACAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN4_3 AAATTACAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
F_SN4_1 AAATTACAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC
Consensus AAATTACAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACAAATGTACAAGTTTCATTGCTGGATTGTGGATGTTGATTGGCAGACAGAATTTGTCACTCAGAGC

1561 1570 1580 1590 1600 1610 1620 1630 1640 1650 1660 1670 1680 1690
|-----|
F_SN4 TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN4_2 TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN4_3 TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA
F_SN4_1 TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA
Consensus TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA

1691 1700 1710 1720 1730 1740 1746
|-----|
F_SN4 TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTAAGATACAATATTGA
F_SN4_2 TCACAATTGAGACTTTGGATGAATGGAGTATGGGCAAGCTAAGATACAATATTGA
F_SN4_3 TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTAAGATACAATAA
F_SN4_1 TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTACTAG
Consensus TCACAATTGAGACTTTGGATGATGGAGTATGGGCAAGCTAAGATACAATAA....

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGTCTTCTCACAAGTTTTTATTCACTTGCATCTCAAATGCTG													
Consensus	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTGTCTTCTCACAAGTTTTTATTCAcTTGCATCTCAAATGCTG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	TAAATGTTCAAAC TGTT CATAGACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCC AATGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAATCCAAACGGGTC													
Consensus	TAAATGTTCAAAC TGTT CATAGACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCC AATGCACCAATGTATTTCAATGGAGTCTAcCATCTATTCTACCAATACAACCCAATGGGTC													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	AGTATGGGGCAATATTGTgTGGGCTCATT CAGTTTCAAGGACTTAATCAATTGGATCAATTTAGA ACCCGCAATTTACCCATCCAACCATTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
Consensus	AGTATGGGGCAATATTGTtTGGGCTCATT CAGTTTCAAGGACTTAATCAATTGGATCAATTTAGA ACCCGCAATTTACCCATCCAACCCTT GATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	CTACCTGGTAACAAGCCTGTTATCTTG TACACTGGAATTGTAGATGCCAACCAACTCAAGTTCAAAC TACGCGATTCCAGCTAACTTATCTGATCCATATCTTCGTGAATGGATCAAGCCTGATAACA													
Consensus	CTACCTGGTAACAAGCCTGTTATCTTG TACACTGGAATTGTAGATGCCAACCAACTCAAGTTCAAAC TACGCGATTCCAGCTAACTTATCTGATCCATATCTTCGTGAATGGATCAAGCCTGATAACA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	ATCCATTGATTGTAGCTGATGCTAGTATCACACAGACCAATTTCTGATCCAACAACCGCATGGATGGGTAAGATGGACATTGGAGATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTAGC													
Consensus	ATCCATTGATTGTAGCTGATGCTAGTATCACACAGACCAATTTCTGATCCAACAACCGCATGGATGGGTAAGATGGACATTGGAGATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTAGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	TATAATGTATAGAACCAAGATTT CATGAATGGGTTAAGGCTAACACCCACTTCAC TCACTAATGGTACTGGAATTTGGGAATGTCC T GATTTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
Consensus	TATAATGTATAGAACCAAGATTT CATGAATGGGTTAAGGCTAACACCCACTTCAC TCACTAATGGTACTGGAATTTGGGAATGTCCcGATTTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAA TAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA													
Consensus	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAA TAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAATGC													
Consensus	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAATGC													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACAATTGGTTCAATGGCCTGTTGAGAATTAGAACCCTAGAACC CAAGGTTCAATTGAGTACAAA													
Consensus	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACAATTGGTTCAATGGCCTGTTGAGAATTAGAACCCTAGAACC CAAGGTTCAATTGAGTACAAA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGA TAAGCAGAGTCATTTGATCCTAGTTGGACTGATATGTATG													
Consensus	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGA TAAGCAGAGTCATTTGATCCTAGTTGGACTGATATGTATG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	CACAGATGTTTGTGGACTTAAGGGTGCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAAC TTAGAAGAAAACACACCTGTTTTCTTCCGAGTTTTCAAGCACAACA													
Consensus	CACAGATGTTTGTGGACTTAAGGGTGCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAAC TTAGAAGAAAACACACCTGTTTTCTTCCGAGTTTTCAAGCACAACA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
F_DN2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_DN2_1	AAATTACAAGGTTCTCTTG TgTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAGC													
Consensus	AAATTACAAGGTTCTCTTG TcTCTGACGCTAAAG													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_TN1	-----													
F_TN1_1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTGTTAGTTTTATTATCCAATAATGTTGTTTTTGCTTCTCATAAAGTTTTTATTCACCTTGCAATCTCAAATGCTG													
Consensus	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTGTTAGTTTTATTATCCAATAATGTTGTTTTTGCTTCTCATAAAGTTTTTATTCACCTTGCAATCTCAAATGCTG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_TN1	-----													
F_TN1_1	TAAATGTTCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCAACGGGTC													
Consensus	TAAATGTTCACTGTTTCATAGAACTGGTTATCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAACCAACGGGTC													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_TN1	-----													
F_TN1_1	AGTATGGGGCAACATTGTTTGGGCTCATTCAAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT													
Consensus	AGTATGGGGCAACATTGTTTGGGCTCATTCAAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCTAACCATTGATCAATTTGGTACATGGTCTGGATCAGCAACTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_TN1	-----													
F_TN1_1	CTACCTGGTAACAACCTGTTATCTTGTAACCTGGAATAGTGGATGCTAACCAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGCCTGATAACA													
Consensus	CTACCTGGTAACAACCTGTTATCTTGTAACCTGGAATAGTGGATGCTAACCAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACATATCTGATCCTTATCTACGTGAATGGATCAGCCTGATAACA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_TN1	-----													
F_TN1_1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAATTTTCGTGATCCACACCCGCTTGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
Consensus	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAATTTTCGTGATCCACACCCGCTTGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_TN1	-----													
F_TN1_1	TATAATGTATAGAACCAAGACTTCATGAATGGGTTAAGGCTAACACCCACTTCCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGGTTGCATTGAAGGAACCTAATGGGCAT													
Consensus	TATAATGTATAGAACCAAGACTTCATGAATGGGTTAAGGCTAACACCCACTTCCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCCGGTTGCATTGAAGGAACCTAATGGGCAT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_TN1	-----													
F_TN1_1	GATCAATATGGTGAAGAACACAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTTCTATTGATA													
Consensus	GATCAATATGGTGAAGAACACAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTACATACCAGATGTTGGTTCTATTGATA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_TN1	-----													
F_TN1_1	GTTGGAAGGGATTGAGATTCGACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAACGC													
Consensus	GTTGGAAGGGATTGAGATTCGACTATGGTAATTTCTATGCATCAAGTCTTTCTATGATCCTAGCAAAATCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAACGC													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_TN1	-----													
F_TN1_1	TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAGTGGTAACCAATTGATTCAATGGCCTGTTGAGAAATTAGAACCCTAGAACCCTAAGGTTCAATTGAGCAACAG													
Consensus	TAAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAGTGGTAACCAATTGATTCAATGGCCTGTTGAGAAATTAGAACCCTAGAACCCTAAGGTTCAATTGAGCAACAG													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
F_TN1	-----													
F_TN1_1	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCCTTTTGCAAGTTTGGAAGGACAGATCATTGATCCTAGTTGGACTGATATGTATG													
Consensus	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCCTTTTGCAAGTTTGGAAGGACAGATCATTGATCCTAGTTGGACTGATATGTATG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
F_TN1	-----													
F_TN1_1	CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTAGAAGAAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAACA													
Consensus	CACAGATGTTTGTGGACTCAAGGTCAGATGTTCAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTAGAAGAAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAACA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
F_TN1	-----													
F_TN1_1	AAATTACAAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACCAATGTACAGAGTTTCATTGCTGGATTTGTGGATGTTGATTGGCAGACAGAATTGTCACTCAGAGC													
Consensus	AAATTACAAGGTTCTCTTGCTCTGATGCTAAAGGTCACCTCTTAAGTTCAATGAACCAATGTACAGAGTTTCATTGCTGGATTTGTGGATGTTGATTGGCAGACAGAATTGTCACTCAGAGC													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
F_TN1	-----													
F_TN1_1	TTGATTGATAATTCAAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACAACGGAACCTGAGCCAA													
Consensus	TTGATTGATAATTCAAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTATCCAACATTGGCGATTAAATGACAGGGCACATTTATTTGCGTTCAACAACGGAACCTGAGCCAA													
	1691	1700	1710	1720	1730	1740	1746							
F_TN1	-----													
F_TN1_1	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA													
Consensus	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_TN2	-----													
F_TN2_1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTTATTATCCAATAATGTTGTTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG													
Consensus	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTTATTATCCAATAATGTTGTTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_TN2	-----													
F_TN2_1	TAAATGTTCAAACGTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAATCCAACGGGTC													
Consensus	TAAATGTTCAAACGTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTATCATCTATTCTACCAATACAATCCAACGGGTC													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_TN2	-----													
F_TN2_1	AGTATGGGGCAATATTGTGTGGGCTCATTCAAGTTTCAAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAACCATTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
Consensus	AGTATGGGGCAATATTGTGTGGGCTCATTCAAGTTTCAAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAACCATTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_TN2	-----													
F_TN2_1	CTACCTGGTAACAAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACTTATCTGATCCATATCTTCGTGAATGGATCAAGCCTGATAATA													
Consensus	CTACCTGGTAACAAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAACCTACGCGGTTCCAGCTAACTTATCTGATCCAATCTTCGTGAATGGATCAAGCCTGATAATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_TN2	-----													
F_TN2_1	ATCCATTGATTGTAGCTGATGCTAGTATCAACAGACCAATTTTCGTGATCCAACAACCGCATGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC													
Consensus	ATCCATTGATTGTAGCTGATGCTAGTATCAACAGACCAATTTTCGTGATCCAACAACCGCATGGATGGGTAAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAAACATAGTAGGGGTTTAGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_TN2	-----													
F_TN2_1	TATAATGTATAGGAGCAAGATTTTCATGAATGGGTAAAGGCTAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGGAATTAATGGGATA													
Consensus	TATAATGTATAGGAGCAAGATTTTCATGAATGGGTAAAGGCTAACACCCACTTCACCTCACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGGAATTAATGGGATA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_TN2	-----													
F_TN2_1	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAAATGGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTACGTACCAGATGTTGGTTCTATTGATA													
Consensus	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTACGTACCAGATGTTGGTTCTATTGATA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_TN2	-----													
F_TN2_1	GTTTGAAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC													
Consensus	GTTTGAAGGGGATTGAGATTTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCTGAGGATGATAATGC													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_TN2	-----													
F_TN2_1	GAAGGGGTGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGTAACAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCTAAGGGTTCAATTGAGTACAG													
Consensus	GAAGGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACCCAGTGGTAACAATTGATTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCTAAGGGTTCAATTGAGTACAG													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
F_TN2	-----													
F_TN2_1	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGAATAAGCAGAGTTATTTGATTCTAGTTGGACTGATATGTATG													
Consensus	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGAATAAGCAGAGTTATTTGATTCTAGTTGGACTGATATGTATG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
F_TN2	-----													
F_TN2_1	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTAGAAGAAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAACA													
Consensus	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTAGAAGAAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAACA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
F_TN2	-----													
F_TN2_1	AAATTACAAGGTTCTCTTGCTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCAGACAGAATTGTCACTCAGAGC													
Consensus	AAATTACAAGGTTCTCTTGCTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCAGACAGAATTGTCACTCAGAGC													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
F_TN2	-----													
F_TN2_1	TTGATTGATAATTCAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCAACATTAGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACAACGGAACCTGAGCCAA													
Consensus	TTGATTGATAATTCAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCAACATTAGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACAACGGAACCTGAGCCAA													
	1691	1700	1710	1720	1730	1740	1746							
F_TN2	-----													
F_TN2_1	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTAA													
Consensus	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAATCACTAG													
	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAcacaAG.....													

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_P18N	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_P18N_1	MDYSSNSRWALPVILVCFEIVLLSNMVVFASHKVFIHLQSQNAVNVQTVHRTGYHFQPEKHWINDPNAPHYFNGVYHLFYQYNPNGSVHGNIVWAHSVSKDLINWINLEPAIYPSKPFDDQFGTWSGSATI													
Consensus	MDYSSNSRWALPVILVCFEIVLLSNMVVFASHKVFIHLQSQNAVNVQTVHRTGYHFQPEKHWINDPNAPHYFNGVYHLFYQYNPNGSVHGNIVWAHSVSKDLINWINLEPAIYPSKPFDDQFGTWSGSATI													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_P18N	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_P18N_1	LPGNKPVILYTGIVDANQTQVQNYAIPANLSDPYLREMIKPDNNPLIVADASINKTKFRDPTTAMHGKDGHWRIYMGSLRKHSRGLAINHYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPVALKGTNGI													
Consensus	LPGNKPVILYTGIVDANQTQVQNYAIPANLSDPYLREMIKPDNNPLIVADASINKTKFRDPTTAMHGKDGHWRIYMGSLRKHSRGLAINHYRSKDFHKWYKAKHPLHSTNGTGNWECPDFFPVALKGTNGI													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_P18N	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_P18N_1	DQYGEEYKYVLKNSHDLTRFEYITLGKYDTKKDRYVPDVGSIKGLRFDYGNFYASKTFYDTSKNRRVIWGSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLVQWPVEELETLRQKVQLSNK													
Consensus	DQYGEEYKYVLKNSHDLTRFEYITLGKYDTKKDRYVPDVGSIKGLRFDYGNFYASKTFYDTSKNRRVIWGSNESDIFPEDDNAKGWAGIQLIPRKVHLDPGKQLVQWPVEELETLRQKVQLSNK													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_P18N	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_P18N_1	KLNGEKVEYTGITPAQADVEYTFASFADKAESFDSSWTDHYAQDYCGLKGADYQGGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKYVLLCSDAKRSTLKFNETHYKVSFAGFYDVLADKKLSLRS													
Consensus	KLNGEKVEYTGITPAQADVEYTFASFADKAESFDSSWTDHYAQDYCGLKGADYQGGGLGPFGLATLATENLEENTPVFFRYFKAQQNYKYVLLCSDAKRSTLKFNETHYKVSFAGFYDVLADKKLSLRS													
	521	530	540	550	560	570	581							
F_P18N	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
F_P18N_1	LIDNSVIESFGAGGKTCITSRYVPTLAINEKAHLFAFNNGTEPITIETLDAWSHGKAKIQY													
Consensus	LIDNSVIESFGAGGKTCITSRYVPTLAINEKAHLFAFNNGTEPITIETLDAWSHGKAKIQY													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_P18N	-----													
F_P18N_1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTTGCTTCTCACAAAGTTTTTATTCATTTGCAATCTCAAATGCTG													
Consensus	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTTATAGTTTATTATCCAATAATGTTGTTTTTGCTTCTCACAAAGTTTTTATTCATTTGCAATCTCAAATGCTG													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_P18N	-----													
F_P18N_1	TAAATGTTCAAACGTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC													
Consensus	TAAATGTTCAAACGTTCATAGAACTGGTTACCATTTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAGTCTACCATCTATTCTACCAATACAACCCAATGGGTC													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_P18N	-----													
F_P18N_1	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGTTTCAAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAACCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
Consensus	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGTTTCAAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAACCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_P18N	-----													
F_P18N_1	CTACCTGGTAACAAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAACCTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA													
Consensus	CTACCTGGTAACAAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAACCTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_P18N	-----													
F_P18N_1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAATTTTCGTGATCCAACAACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC													
Consensus	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAATTTTCGTGATCCAACAACCGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGGAACATAGTAGGGGTTTAGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_P18N	-----													
F_P18N_1	TATAATGTATAGAAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACCTCACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTTCCCTGTTGCATTGAAGGGAATTAATGGGATA													
Consensus	TATAATGTATAGAAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACCTCACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTTCCCTGTTGCATTGAAGGGAATTAATGGGATA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_P18N	-----													
F_P18N_1	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA													
Consensus	GATCAATATGGTGAAGAATATAAATATGTGCTTAAGAAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_P18N	-----													
F_P18N_1	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAATGC													
Consensus	GTTGGAAGGGATTGAGATTCGATTATGGTAATTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATTCCCTGAGGATGATAATGC													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_P18N	-----													
F_P18N_1	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCTAAGGTTCAATTGAGTACAAA													
Consensus	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGATCCAAGTGGTAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCTAAGGTTCAATTGAGTACAAA													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
F_P18N	-----													
F_P18N_1	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTTTCTTTTGCAAGTTTGGAATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG													
Consensus	AAGTTGAACAATGGTGAAGAGGTTGAAGTTACAGGAATCACACCTGCACAGGCAGATGTTGAAGTGACATTTTCTTTTGCAAGTTTGGAATAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
F_P18N	-----													
F_P18N_1	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTGGGAAGAAAACACACCTGTTTTCTTCCGAGTTTTCAAGCACAACA													
Consensus	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAAACCTGGGAAGAAAACACACCTGTTTTCTTCCGAGTTTTCAAGCACAACA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
F_P18N	-----													
F_P18N_1	AAATTACAAGGTTCTTTTGCTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAGC													
Consensus	AAATTACAAGGTTCTTTTGCTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGGCTGACAAGAATTGTCACTCAGAGC													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
F_P18N	-----													
F_P18N_1	TTGATTGATAATTCAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACAATGGAAGTGAAGC													
Consensus	TTGATTGATAATTCAGTTATAGAAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTATCCAACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACAATGGAAGTGAAGC													
	1691	1700	1710	1720	1730	1740	1746							
F_P18N	-----													
F_P18N_1	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA													
Consensus	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
F_P40N2	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG														
F_P40N2_2	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG														
F_P40N2_1	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG														
Consensus	ATGGATTATTCATCTAATTCCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCTTTTTATAGTTTTATTATCCAATAATGTTGTTTTGCTTCTCACAAAGTTTTATTCACTTGCAATCTCAAATGCTG														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
F_P40N2	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAATTTACCATCTATTCTACCAATACAACCCAAACGGGGTC														
F_P40N2_2	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAATTTACCATCTATTCTACCAATACAACCCAAACGGGGTC														
F_P40N2_1	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAATTTACCATCTATTCTACCAATACAACCCAAATGGGGTC														
Consensus	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTTCAGCCAGAAAAACATTGGATCAATGATCCCAATGCACCAATGTATTTCAATGGAATTTACCATCTATTCTACCAATACAACCCAAACGGGGTC														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
F_P40N2	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCATTGATCAATTTGGTACGTGGTCTGGATCTGCAACTATT														
F_P40N2_2	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCATTGATCAATTTGGTACGTGGTCTGGATCTGCAACTATT														
F_P40N2_1	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCCTTTGATCAATTCGGTACGTGGTCTGGATCTGCAACTATT														
Consensus	AGTATGGGGCAATATTGTTTGGGCTCATTTCAGTTTCAAGGACTTAATCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAARCCaTTTATGATCAATTTGGTACGTGGTCTGGATCTGCAACTATT														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
F_P40N2	CTACCTGGTAACAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAAACTACGCGATTCCAGCTAATTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
F_P40N2_2	CTACCTGGTAACAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAAACTACGCGATTCCAGCTAATTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
F_P40N2_1	CTACCTGGTAACAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAAACTACGCGATTCCAGCTAATTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
Consensus	CTACCTGGTAACAGCCTGTTATCTTGTAACCTGGAATAGTAGATGCCAACCAACTCAAGTTCAAAACTACGCGATTCCAGCTAATTATCTGATCCATATCTCCGTGAATGGATCAAGCCTGATAACA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
F_P40N2	ATCCATTGATTGTAGCTGATGATAGTATCAACAAGACCAATTTTCGTGATCCACAACTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC														
F_P40N2_2	ATCCATTGATTGTAGCTGATGATAGTATCAACAAGACCAATTTTCGTGATCCACAACTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC														
F_P40N2_1	ATCCATTGATTGTAGCTGATGATAGTATCAACAAGACCAATTTTCGTGATCCACAACTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC														
Consensus	ATCCATTGATTGTAGCTGATGATAGTATCAACAAGACCAATTTTCGTGATCCACAACTGCATGGATGGGTAAGATGGACATTGGAGAATTGTGATGGGAAGTTTGAGAAACATAGTAGGGGTTTAGC														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
F_P40N2	AATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACTCAACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCTGTTGCATTGAAGGGAAGTAATGGGATA														
F_P40N2_2	AATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACTCAACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCTGTTGCATTGAAGGGAAGTAATGGGATA														
F_P40N2_1	AATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACTCAACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCTGTTGCATTGAAGGGAAGTAATGGGATA														
Consensus	AATAATGTATAGAGCAAGATTTTCATGAATGGGTTAAGGCTAACACCCACTTCACTCAACTAACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCTGTTGCATTGAAGGGAAGTAATGGGATA														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
F_P40N2	GATCAATATGATGAAGAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
F_P40N2_2	GATCAATATGATGAAGAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
F_P40N2_1	GATCAATATGATGAAGAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
Consensus	GATCAATATGATGAAGAATATAAATATGTGCTTAAGAATAGTATGGATCTTACTCGATTTGAGTACTATACTCTTGGTAGTACGATACAAAAAAGATAGGTATGTTCCAGATGTTGGTTCTATTGATA														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
F_P40N2	GTTTGAAGGGGATTGAGATTCGATTATGGTAACCTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGGATGATAATGC														
F_P40N2_2	GTTTGAAGGGGATTGAGATTCGATTATGGTAACCTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGGATGATAATGC														
F_P40N2_1	GTTTGAAGGGGATTGAGATTCGATTATGGTAACCTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGGATGATAATGC														
Consensus	GTTTGAAGGGGATTGAGATTCGATTATGGTAACCTTCTATGCATCAAGACTTTCTATGATACTAGCAAAAACCGAAGGGTTATTTGGGGTTGGTCTAATGAATCAGATATATCCCTGAGGATGATAATGC														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
F_P40N2	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACGCAAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCAAAGGTTCAATTGAGTAACAAA														
F_P40N2_2	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACGCAAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCAAAGGTTCAATTGAGTAACAAA														
F_P40N2_1	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACGCAAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCAAAGGTTCAATTGAGTAACAAA														
Consensus	GAAGGATGGGCTGGAATTCATTGATCCACGTAAGTATGGCTTGACGCAAGTGGAACAATTGGTTCAATGGCCTGTTGAAGAATTAGAACCCTAGAACCCAAAGGTTCAATTGAGTAACAAA														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
F_P40N2	AAGTTGAACAATGGTGAAAAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
F_P40N2_2	AAGTTGAACAATGGTGAAAAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
F_P40N2_1	AAGTTGAACAATGGTGAAAAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
Consensus	AAGTTGAACAATGGTGAAAAGGTTGAAGTTACAGGGATCACACCTGCACAGGCAGATGTTGAAGTGACATTCTCTTTGCAAGTTTGGATAAAGCAGAGTCATTTGATTCTAGTTGGACTGATATGTATG														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
F_P40N2	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACTTAGAAGAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAAACA														
F_P40N2_2	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACTTAGAAGAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAAACA														
F_P40N2_1	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACTTAGAAGAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAAACA														
Consensus	CACAGATGTTTGTGGACTCAAGGGTGCAGATGTACAAGGTGGGCTTGGGCCATTTGGTCTTGCTACATTGGCTACTGAAACTTAGAAGAAACACACCTGTTTTCTCCGAGTTTTCAAGCACAAACA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
F_P40N2	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGACAGACAAGAATTGTCCTCAGAGC														
F_P40N2_2	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGACAGACAAGAATTGTCCTCAGAGC														
F_P40N2_1	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGACAGACAAGAATTGTCCTCAGAGC														
Consensus	AAATTACAAGGTTCTCTTGTTCTGACGCTAAAGGTCACCTCTTAAGTTCAATGAACAATGTACAAGTTTCATTTGCTGGATTTGTGGATGTTGATTTGACAGACAAGAATTGTCCTCAGAGC														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
F_P40N2	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA														
F_P40N2_2	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA														
F_P40N2_1	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA														
Consensus	TTGATTGATAATTCAGTTATAGAAGTTTTGGTGCTGGTGGAAAGACATGTATAACATCGAGGGTTTATCCACATTGGCGATTAAATGAGAAGGCACATTTATTTGCGTTCAACACGGAACTGAGCCAA														
	1691	1700	1710	1720	1730	1740	1746								
F_P40N2	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA														
F_P40N2_2	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA														
F_P40N2_1	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA														
Consensus	TCACAATTGAGACTTTGGATGCATGGAGTATGGGCAAGCTAAGATACAATATTGA														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_P54N1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
F_P54N1_3	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
F_P54N1_5	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
F_P54N1_1	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
F_P54N1_2	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
F_P54N1_4	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
Consensus	ATGGATTATTCATCTAATTCTCGTTGGGCTTTGCCAGTTATCTTAGTTTGCCTTTTTATAGTTTTATTATCCAAATAATGTTGTTTTGCTTCTCACAAAGT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_P54N1	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
F_P54N1_3	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
F_P54N1_5	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
F_P54N1_1	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
F_P54N1_2	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
F_P54N1_4	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
Consensus	TAAATGTTCAAACCTGTTTCATAGAACTGGTTACCATTTCAGCCAGAAAACATTGGATCAATGATCCCAATGCACCAATG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_P54N1	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
F_P54N1_3	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
F_P54N1_5	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
F_P54N1_1	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
F_P54N1_2	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
F_P54N1_4	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
Consensus	AGTATGGGGCAATATTGTTTGGGCTCATTCAAGGACTTAACTCAATTGGATCAATTTAGAACCCGCAATTTACCCATCCAAACCCCTTGGATCAATTCGGTACGTGGTCTGGATCTGCACCTATT													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_P54N1	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
F_P54N1_3	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
F_P54N1_5	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
F_P54N1_1	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
F_P54N1_2	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
F_P54N1_4	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
Consensus	CTACCTGGTAACAGCCTGTTATCTTGACACTGGATAGTAGATGCCAACCAACTCAGTTCAAACACTACGCGATTCCAGCTAACTTATCTGATCCATATCTCCGTAATGGATCAGCCTGATACCA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_P54N1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
F_P54N1_3	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
F_P54N1_5	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
F_P54N1_1	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
F_P54N1_2	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
F_P54N1_4	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
Consensus	ATCCATTGATTGTAGCTGATGCTAGCATCAACAGACCAAAATTCCTGATCCAAACCCGCAATGGATGGGTAAGATGGACATTGGAGAAATTTGGATGGGAGTTTGGAGAAACATAGTAGGGGTTTACG													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_P54N1	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
F_P54N1_3	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
F_P54N1_5	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
F_P54N1_1	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
F_P54N1_2	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
F_P54N1_4	TATAATGTATAGAGCAAGATTTTCATGAATGGTTAAGGCTAAACACCCACTTCACTCAACTAATGGTACTGGAACCTGGGAATGTCCTGATTTTTCCCTGTTGCATTGAAGGAACTAATGGGATA													
Consensus	TATAATGTATAGAGCAAGATTT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	AT	G	G	A	T	T	C	A	T	T	C	T	C	G
F_P54N2_3	AT	G	G	A	T	T	C	A	T	T	C	T	C	G
F_P54N2_1	AT	G	G	A	T	T	C	A	T	T	C	T	C	G
F_P54N2_2	AT	G	G	A	T	T	C	A	T	T	C	T	C	G
Consensus	AT	G	G	A	T	T	C	A	T	T	C	T	C	G
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P54N2_3	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P54N2_1	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P54N2_2	T	A	A	T	G	T	T	C	A	A	A	C	T	T
Consensus	T	A	A	T	G	T	T	C	A	A	A	C	T	T
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P54N2_3	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P54N2_1	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P54N2_2	A	G	T	A	T	G	G	G	C	A	A	T	T	G
Consensus	A	G	T	A	T	G	G	G	C	A	A	T	T	G
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	C	T	A	C	T	G	G	T	A	C	A	A	A	C
F_P54N2_3	C	T	A	C	T	G	G	T	A	C	A	A	A	C
F_P54N2_1	C	T	A	C	T	G	G	T	A	C	A	A	A	C
F_P54N2_2	C	T	A	C	T	G	G	T	A	C	A	A	A	C
Consensus	C	T	A	C	T	G	G	T	A	C	A	A	A	C
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	A	T	C	C	A	T	T	G	A	T	T	G	A	T
F_P54N2_3	A	T	C	C	A	T	T	G	A	T	T	G	A	T
F_P54N2_1	A	T	C	C	A	T	T	G	A	T	T	G	A	T
F_P54N2_2	A	T	C	C	A	T	T	G	A	T	T	G	A	T
Consensus	A	T	C	C	A	T	T	G	A	T	T	G	A	T
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	T	A	T	A	T	G	T	A	G	A	C	A	A	A
F_P54N2_3	T	A	T	A	T	G	T	A	G	A	C	A	A	A
F_P54N2_1	T	A	T	A	T	G	T	A	G	A	C	A	A	A
F_P54N2_2	T	A	T	A	T	G	T	A	G	A	C	A	A	A
Consensus	T	A	T	A	T	G	T	A	G	A	C	A	A	A
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	G	A	T	C	A	A	T	A	T	A	A	T	A	T
F_P54N2_3	G	A	T	C	A	A	T	A	T	A	A	T	A	T
F_P54N2_1	G	A	T	C	A	A	T	A	T	A	A	T	A	T
F_P54N2_2	G	A	T	C	A	A	T	A	T	A	A	T	A	T
Consensus	G	A	T	C	A	A	T	A	T	A	A	T	A	T
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	G	T	T	G	A	G	G	G	A	T	T	C	A	T
F_P54N2_3	G	T	T	G	A	G	G	G	A	T	T	C	A	T
F_P54N2_1	G	T	T	G	A	G	G	G	A	T	T	C	A	T
F_P54N2_2	G	T	T	G	A	G	G	G	A	T	T	C	A	T
Consensus	G	T	T	G	A	G	G	G	A	T	T	C	A	T
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	G	A	A	G	G	A	T	T	C	A	A	T	T	C
F_P54N2_3	G	A	A	G	G	A	T	T	C	A	A	T	T	C
F_P54N2_1	G	A	A	G	G	A	T	T	C	A	A	T	T	C
F_P54N2_2	G	A	A	G	G	A	T	T	C	A	A	T	T	C
Consensus	G	A	A	G	G	A	T	T	C	A	A	T	T	C
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	A	A	G	T	T	G	A	A	G	G	T	T	G	A
F_P54N2_3	A	A	G	T	T	G	A	A	G	G	T	T	G	A
F_P54N2_1	A	A	G	T	T	G	A	A	G	G	T	T	G	A
F_P54N2_2	A	A	G	T	T	G	A	A	G	G	T	T	G	A
Consensus	A	A	G	T	T	G	A	A	G	G	T	T	G	A
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	C	A	C	A	G	A	T	G	T	T	G	G	A	C
F_P54N2_3	C	A	C	A	G	A	T	G	T	T	G	G	A	C
F_P54N2_1	C	A	C	A	G	A	T	G	T	T	G	G	A	C
F_P54N2_2	C	A	C	A	G	A	T	G	T	T	G	G	A	C
Consensus	C	A	C	A	G	A	T	G	T	T	G	G	A	C
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	A	A	A	T	T	C	A	A	G	G	T	T	C	T
F_P54N2_3	A	A	A	T	T	C	A	A	G	G	T	T	C	T
F_P54N2_1	A	A	A	T	T	C	A	A	G	G	T	T	C	T
F_P54N2_2	A	A	A	T	T	C	A	A	G	G	T	T	C	T
Consensus	A	A	A	T	T	C	A	A	G	G	T	T	C	T
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
		-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	
F_P54N2	T	T	G	A	T	T	G	A	A	G	T	T	T	G
F_P54N2_3	T	T	G	A	T	T	G	A	A	G	T	T	T	G
F_P54N2_1	T	T	G	A	T	T	G	A	A	G	T	T	T	G
F_P54N2_2	T	T	G	A	T	T	G	A	A	G	T	T	T	G
Consensus	T	T	G	A	T	T	G	A	A	G	T	T	T	G
	1691	1700	1710	1720	1730	1740	1746							
		-----	-----	-----	-----	-----	-----							
F_P54N2	T	C	A	A	T	T	G	A	C	A	T	T	T	G
F_P54N2_3	T	C	A	A	T	T	G	A	C	A	T	T	T	G
F_P54N2_1	T	C	A	A	T	T	G	A	C	A	T	T	T	G
F_P54N2_2	T	C	A	A	T	T	G	A	C	A	T	T	T	G
Consensus	T	C	A	A	T	T	G	A	C	A	T	T	T	G

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
F_P18N	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
F_P54N1	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
F_P40N1	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
F_P54N2	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
F_P40N2	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
Consensus	AT	G	G	A	T	A	T	T	C	A	A	T	T	C
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
F_P18N	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P54N1	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P40N1	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P54N2	T	A	A	T	G	T	T	C	A	A	A	C	T	T
F_P40N2	T	A	A	T	G	T	T	C	A	A	A	C	T	T
Consensus	T	A	A	T	G	T	T	C	A	A	A	C	T	T
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
F_P18N	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P54N1	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P40N1	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P54N2	A	G	T	A	T	G	G	G	C	A	A	T	T	G
F_P40N2	A	G	T	A	T	G	G	G	C	A	A	T	T	G
Consensus	A	G	T	A	T	G	G	G	C	A	A	T	T	G
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
F_P18N	C	T	A	C	T	G	G	T	A	A	C	A	A	C
F_P54N1	C	T	A	C	T	G	G	T	A	A	C	A	A	C
F_P40N1	C	T	A	C	T	G	G	T	A	A	C	A	A	C
F_P54N2	C	T	A	C	T	G	G	T	A	A	C	A	A	C
F_P40N2	C	T	A	C	T	G	G	T	A	A	C	A	A	C
Consensus	C	T	A	C	T	G	G	T	A	A	C	A	A	C
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
F_P18N	A	T	C	A	T	T	G	A	T	T	G	A	T	T
F_P54N1	A	T	C	A	T	T	G	A	T	T	G	A	T	T
F_P40N1	A	T	C	A	T	T	G	A	T	T	G	A	T	T
F_P54N2	A	T	C	A	T	T	G	A	T	T	G	A	T	T
F_P40N2	A	T	C	A	T	T	G	A	T	T	G	A	T	T
Consensus	A	T	C	A	T	T	G	A	T	T	G	A	T	T
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
F_P18N	T	A	T	A	T	G	T	A	A	C	A	A	A	A
F_P54N1	T	A	T	A	T	G	T	A	A	C	A	A	A	A
F_P40N1	T	A	T	A	T	G	T	A	A	C	A	A	A	A
F_P54N2	T	A	T	A	T	G	T	A	A	C	A	A	A	A
F_P40N2	T	A	T	A	T	G	T	A	A	C	A	A	A	A
Consensus	T	A	T	A	T	G	T	A	A	C	A	A	A	A
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
F_P18N	G	A	T	C	A	A	T	A	T	G	T	G	C	T
F_P54N1	G	A	T	C	A	A	T	A	T	G	T	G	C	T
F_P40N1	G	A	T	C	A	A	T	A	T	G	T	G	C	T
F_P54N2	G	A	T	C	A	A	T	A	T	G	T	G	C	T
F_P40N2	G	A	T	C	A	A	T	A	T	G	T	G	C	T
Consensus	G	A	T	C	A	A	T	A	T	G	T	G	C	T
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
F_P18N	G	T	T	G	G	A	G	G	A	T	T	G	A	T
F_P54N1	G	T	T	G	G	A	G	G	A	T	T	G	A	T
F_P40N1	G	T	T	G	G	A	G	G	A	T	T	G	A	T
F_P54N2	G	T	T	G	G	A	G	G	A	T	T	G	A	T
F_P40N2	G	T	T	G	G	A	G	G	A	T	T	G	A	T
Consensus	G	T	T	G	G	A	G	G	A	T	T	G	A	T
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
F_P18N	G	A	A	G	G	A	T	T	G	A	A	T	T	G
F_P54N1	G	A	A	G	G	A	T	T	G	A	A	T	T	G
F_P40N1	G	A	A	G	G	A	T	T	G	A	A	T	T	G
F_P54N2	G	A	A	G	G	A	T	T	G	A	A	T	T	G
F_P40N2	G	A	A	G	G	A	T	T	G	A	A	T	T	G
Consensus	G	A	A	G	G	A	T	T	G	A	A	T	T	G
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
F_P18N	A	A	G	T	T	G	A	A	A	G	G	T	T	G
F_P54N1	A	A	G	T	T	G	A	A	A	G	G	T	T	G
F_P40N1	A	A	G	T	T	G	A	A	A	G	G	T	T	G
F_P54N2	A	A	G	T	T	G	A	A	A	G	G	T	T	G
F_P40N2	A	A	G	T	T	G	A	A	A	G	G	T	T	G
Consensus	A	A	G	T	T	G	A	A	A	G	G	T	T	G
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
F_P18N	C	A	C	A	A	G	A	T	T	G	T	T	G	A
F_P54N1	C	A	C	A	A	G	A	T	T	G	T	T	G	A
F_P40N1	C	A	C	A	A	G	A	T	T	G	T	T	G	A
F_P54N2	C	A	C	A	A	G	A	T	T	G	T	T	G	A
F_P40N2	C	A	C	A	A	G	A	T	T	G	T	T	G	A
Consensus	C	A	C	A	A	G	A	T	T	G	T	T	G	A
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
F_P18N	A	A	A	T	T	A	C	A	A	G	T	T	T	A
F_P54N1	A	A	A	T	T	A	C	A	A	G	T	T	T	A
F_P40N1	A	A	A	T	T	A	C	A	A	G	T	T	T	A
F_P54N2	A	A	A	T	T	A	C	A	A	G	T	T	T	A
F_P40N2	A	A	A	T	T	A	C	A	A	G	T	T	T	A
Consensus	A	A	A	T	T	A	C	A	A	G	T	T	T	A
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
F_P18N	T	T	G	A	T	T	G	A	A	A	G	G	T	T
F_P54N1	T	T	G	A	T	T	G	A	A	A	G	G	T	T
F_P40N1	T	T	G	A	T	T	G	A	A	A	G	G	T	T
F_P54N2	T	T	G	A	T	T	G	A	A	A	G	G	T	T
F_P40N2	T	T	G	A	T	T	G	A	A	A	G	G	T	T
Consensus	T	T	G	A	T	T	G	A	A	A	G	G	T	T
	1691	1700	1710	1720	1730	1740	1746							
F_P18N	T	C	A	A	T	T	G	A	C	T	T	T	T	G
F_P54N1	T	C	A	A	T	T	G	A	C	T	T	T	T	G
F_P40N1	T	C	A	A	T	T	G	A	C	T	T	T	T	G
F_P54N2	T	C	A	A	T	T	G	A	C	T	T	T	T	G
F_P40N2	T	C	A	A	T	T	G	A	C	T	T	T	T	G
Consensus	T	C	A	A	T	T	G	A	C	T	T	T	T	G

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_SA	MELFMKSSSLWGLEIYLF CFFIVLSNINKVFGSHNVFLDLQSSSAISVKNVHRTGFHFQPPKHWINDPNAPHYYNGVYHLFYQYNPKGSVMGNIYWAHSVSKDLINWHLLEPAIYPSKKFDKYGAWSGSA														
F_SN4	MD-Y-SSNSRWALPVILVCFVLLSNMVVFAHSHKVF IHLQSQNAYNVHTVHRTGYHFQPEKHWINDPNAPHYFNGVYHLFYQYNPNKGSVMGNIYWAHSVSKDLINWHLLEPAIYPSKPFQDFGTWSGSA														
Consensus	M#.%.kSnSlMaLeiilfCFFiVLlninkVFasHkVFidLQsqnAinVhnVHRTG%HFQPeKHWINDPNAPHY%NGVYHLFYQYNPKGSVMGNIYWAHSVSKDLINWHLLEPAIYPSKkFDk%GaWSGSA														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_SA	TILPNMKPVILYTGIVVDSHDTQVQNYAIPANLSDPFLRKWVKPNNPLIIPDNSINKTKFRDPTTAWMGVYDGVWRIYIGSMRKH-RGMALLYRSRDFIKWYKAQHPLHSSPHTGNWECPDFFFPVSLNNTN														
F_SN4	TILPGMKPVILYTGIVDANQTQVQNYAVPANISDPYLRWIKPDNNPLIYADASINKTKFRDPTTAWMGKDGHWRIYMGSLRKHSRGLAIMYRSKDFMKWYKAKHPLHSTNGTNWECPDFFFPVALKGTN														
Consensus	TILPgMKPVILYTGIVDah#TQVQNYAiPANiSDP%LRWiKP#NNPLIiaDaSINKTKFRDPTTAWMGkDghWRIYiGS\$RKH.RG\$Ai\$YRSkDFiKWYKakHPLHSsngTNWECPDFFFPVaLkgTN														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_SA	GLDASVYRGKNVYKYLKNSLDVNRFDYYTIGMYDTRKDRYIPDNNSIDGCKGLRLDYGNFYASKSFYDPTKNRRIYVWGMTNESDVLPDDEIKKGWAGIQAIIPRKVWLDHSGKQLIQWPIEELETLRKQKIQ														
F_SN4	GQDQ-Y-GEEHKYVLKNSMDLTRFEYYTLGKYDTKKDRYIPDVGSIDSMKGLRFDYGNFYASKSFYDPSKNRRIYVWGMTNESDIFPEDDIAKGWAGIQLIPRKVWLDPSGKQLIQWPVEELETLRKQKYQ														
Consensus	GIda.Y.Ge#hKYVLKNS\$DlnRF#YYTiGkYDTkKDRYIPDngSIDgcKGLRfDYGNFYASKSFYDPSKNRRIiWGMsNESDifP#D#iAKGWAGIQaIPRKVWLDhSGKQLIQWpiEELETLRkQKiQ														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
E_SA	LNNKKLSKGEMFEVKGISASQSDIEVSFSFSSLNKAEQFDPNWADLYAQDVCaIKGSTIQGGLGPFGLATLASKNLEEYTPVFFRVFKAQKNYKVLHCSDARRSTHRQNEAMYPKPSFAGYVDYDLVDMKK														
F_SN4	LSNKKLNNGEKVEVTGITPAQADVEYTFSFASLDKAESFDPSTWTHYAQDVCGLKGADYQGGLGPFGLATLATENLEENTPVFFRVFKAQQNYKVLCSDAKRSTLKFNETHYKYVSFAGFYVDYDLAD-KK														
Consensus	LnNKKLnkGEkfEVkGI\$aaQaDiEv\$FSFaSL#KAeqFDPnWaD\$YAQDVCaIKGadiQGGLGPFGLATLaseNLEENTPVFFRVFKAQkNYKVL\$CSDAkRST\$kfNEaMYKpSFAG%VDYDLAd.KK														
	521	530	540	550	560	570	580	586							
	-----+-----+-----+-----+-----+-----+-----+-----														
E_SA	LSLRSLIDNSVYESFGAGGKTCITSRVYPTLAIHDNAHLFVFNNGSETITITETLNAWSHMDVPKMH														
F_SN4	LSLRSLIDNSVIESFGAGGKTCITSRVYPTLAINDKAHLFAFNNGTETITITETLDNAWSHGAKIKY														
Consensus	LSLRSLIDNSViESFGAGGKTCITSRVYPTLAIhdkAHLfAfNNGsEpITITETL#AWSHdkaKih.														

Appendix 3.3

A sequence was defined as an allele when it was found twice in two independent PCRs. Additionally, the consensus sequence of all alleles found in one genotype was used for allele definition when variable sequence polymorphisms occurred.

In the following Table, *pCD111* and *pCD141* alleles are listed, which do not correspond to this definition because only one full-length sequence was obtained. Nevertheless, for some of the listed alleles independent PCR amplifications gave rise to sequences that after partial sequencing indicated the existence of the alleles as mentioned above, clearly implying their existence. These additional sequences exhibited frame shifts or other modifications (e.g. missing or modified start and stop codons) and, therefore, were not completely sequenced. Cloning of *pCD111* and *pCD141* alleles was carried out in addition to *Pain-1*, *invGE*, and *invGF* allele cloning, which were in the focus of the present study.

Table A3.3.1: Overview of fully and partially sequenced *pCD111* and *pCD141* alleles.

Gene	Full-length allele	Additional partial sequences
<i>pCD111</i>	<i>pCD111_S3</i>	No additional sequences
	<i>pCD111_D1</i>	No additional sequences
	<i>pCD111_T1</i>	No additional sequences
	<i>pCD111_P40_1</i>	2
	<i>pCD111_P54_2</i>	No additional sequences
<i>pCD141</i>	<i>pCD141_S1</i>	2
	<i>pCD141_D2</i>	No additional sequences
	<i>pCD141_T3</i>	No additional sequences

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_S1	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_S1_1	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_S3	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_S2	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_S2_1	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
Consensus	MDCLKKSSLFSLPIFLLYFSIILSFNNGVNAHSHKVFPGLQSTSTVDVKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQNPYPGSMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_S1	ILPNNKPIILYTGIVDAKNTQVQNYAIPANISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSKNFHKTQIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_S1_1	ILPNNKPIILYTGIVDAKNTQVQNYAIPANISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSKNFHKTQIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_S3	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSKNFHKTQIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_S2	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSKNFHKTQIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_S2_1	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSKNFHKTQIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
Consensus	ILPNNKPIILYTGIVDAKNTQVQNYAIPA#ISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGQDGYWRTLIGSYWgKQGLAILYKSK#FHKWTK!QHPLHSVDGTGNWECPDFFPVLLHGTNGL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_S1	DASYNKKNIKHALKVSLDVTRFEYYTVGIDYDTKKDRYIPDKTSIDGwKGLRLDYGNYYASKSFYDPSKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
CD111_S1_1	DASYNKKNIKHALKVSLDVTRFEYYTVGIDYDTKKDRYIPDKTSIDGwKGLRLDYGNYYASKSFYDPSKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
CD111_S3	DASYNKKNIKHVLLKVSVDVTRFEYYTVGIDYDTKKDRYIPDKTSIDGwKGLRLDYGNYYASKSFYDPSKNRRIMHGWANESDTYNDDIKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
CD111_S2	DASYNKKNIKHALKVSLDVTRFEYYTVGKYDTKKDRYIPDKTSIDGLNGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
CD111_S2_1	DASYNKKNIKHALKVSLDVTRFEYYTVGKYDTKKDRYIPDKTSIDGLNGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
Consensus	DASYNKKNIKHALKVSLDVTRFEYYTVGiYDTKKDRYIPDKTSIDGwKGLRLDYGNYYASKSFYDpsKNRRIMHGWANESDTYNDD!KKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIREQKVQLSN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_S1	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
CD111_S1_1	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
CD111_S3	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
CD111_S2	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
CD111_S2_1	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
Consensus	RKLKKGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPWMDNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRVFKAQDKYKVLHCSASRSTLKNDKTHYKPSFAGYVDYDLINKTSLR													
	521	530	540	550	560	570	580	589						
CD111_S1	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_S1_1	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSNH													
CD111_S3	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_S2	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_S2_1	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
Consensus	SLIDHSVYESFGAGGKTCITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSy													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_T1	MDCLKKSSLFSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPQNWINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_T2	MDCLKKSSLFSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNWNINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_T2_2	MDCLKKSSLFSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNWNINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_T2_1	MDCLKKSSLFSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNWNINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
Consensus	MDCLKKSSLFSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPP#NWINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYMAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_T1	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDQGYWRTLIGSYWgKQGLAILYKSKNLHKWTKIQHPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_T2	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDQGYWRTLIGSYWgKKGLAILYKSRDFHKWTKYQDPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_T2_2	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDQGYWRTLIGSYWgKKGLAILYKSRDFHKWTKYQDPLHSVDGTGNWECPDFFPVLLHGTNGL													
CD111_T2_1	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDQGYWRTLIGSYWgKKGLAILYKSRDFHKWTKYQDPLHSVDGTGNWECPDFFPVLLHGTNGL													
Consensus	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDQGYWRTLIGSYWgKkGLAILYKSR#fHKWTK!QdPLHSVDGTGNWECPDFFPVLLHGTNGL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_T1	DASYNKKNIKHVgLVSLDVTREFEYTYVGQYDTKKDRYIPDKTSIDGWgKGLRLDYGNYYASKSFYDPSkNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETREQKVQLSN													
CD111_T2	DASYNKKNIKHgLVSLDVTREFEYTYVGbYDTKKDRYIPDKTSIDGWgGLRLDYGNYYASKSFYDLRkNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETREQKVQLSN													
CD111_T2_2	DASYNKKNIKHgLVSLDVTREFEYTYVGbYDTKKDRYIPDKTSIDGWgGLRLDYGNYYASKSFYDLRkNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETREQKVQLSN													
CD111_T2_1	DASYNKKNIKHgLVSLDVTREFEYTYVGbYDTKKDRYIPDKTSIDGWgGLRLDYGNYYASKSFYDLRkNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETREQKVQLSN													
Consensus	DASYNKKNIKHgLVSLDVTREFEYTYVGbYDTKKDRYIPDKTSIDGWgGLRLDYGNYYASKSFYDLrkNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETREQKVQLSN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_T1	RKLNKGDKIEYKGITPAQADYEYTFSTSLDKAETFDPNWgNLYAQDYCAIKGSgVQGGLGPFGLLTLASQNLEEYTPVFFRYFKAQDKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDYDLTNKTLSLR													
CD111_T2	RKLNKGDKIEYKGITPAQADYEYTFSSSLDKAEPFDPNWgNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRYFKAQNKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDYDLTNKTLSLR													
CD111_T2_2	RKLNKGDKIEYKGITPAQADYEYTFSSSLDKAEPFDPNWgNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRYFKAQNKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDYDLTNKTLSLR													
CD111_T2_1	RKLNKGDKIEYKGITPAQADYEYTFSSSLDKAEPFDPNWgNLYAQDYCAIKGSTVQGGLGPFGLLTLASQNLEEYTPVFFRYFKAQNKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDYDLTNKTLSLR													
Consensus	RKLNKGDKIEYKGITPAQADYEYTFSSSLDKAEPFDPNWgNLYAQDYCAIKGSgVQGGLGPFGLLTLASQNLEEYTPVFFRYFKAQ#KYKVLHCSDASRSTLKNDKTHYKPSFAGYVDYDLTNKTLSLR													
	521	530	540	550	560	570	580	589						
CD111_T1	SLIDHSVYESFGAGGKTcITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_T2	SLIDHSVYESFGAGGKTcITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_T2_2	SLIDHSVYESFGAGGKTcITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
CD111_T2_1	SLIDHSVYESFGAGGKTRITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													
Consensus	SLIDHSVYESFGAGGKTcITSRYYP TLAIYDNAHLFVFNNGTETIKIKSLNAWTHGKPKMNWSFGHSSY													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_S1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1_1	ATGGATTGTTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTTGCAATCTACAAGCA													
Consensus	ATGGATTGTTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTTGCAATCTACAAGCA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAAAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
CD111_S1_1	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAAAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
Consensus	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAAAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTTGACAATATGGTACATGGTCCGGGTCAGCCACA													
CD111_S1_1	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTTGACAATATGGTACATGGTCCGGGTCAGCCACA													
Consensus	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTTGACAATATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	ATCTTACCCAACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAGTCCAAACTATGCAATCCAGCCACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
CD111_S1_1	ATCTTACCCAACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAGTCCAAACTATGCAATCCAGCCACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
Consensus	ATCTTACCCAACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAGTCCAAACTATGCAATCCAGCCACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGCGACCCAACACATGTTGGTTGGGTGAGGATGGTATTGGAGAACCTTGATAGGGAGTGTGTGGGGAAACAGGATTGGC													
CD111_S1_1	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGCGACCCAACACATGTTGGTTGGGTGAGGATGGTATTGGAGAACCTTGATAGGGAGTGTGTGGGGAAACAGGATTGGC													
Consensus	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGCGACCCAACACATGTTGGTTGGGTGAGGATGGTATTGGAGAACCTTGATAGGGAGTGTGTGGGGAAACAGGATTGGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	AATATTGTATAAAGTAAAAATTTTCATGAATGGACCAAGATTCAACATCCACTTCATTAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
CD111_S1_1	AATATTGTATAAAGTAAAAATTTTCATGAATGGACCAAGATTCAACATCCACTTCATTAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
Consensus	AATATTGTATAAAGTAAAAATTTTCATGAATGGACCAAGATTCAACATCCACTTCATTAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	GATGCCTCATACACAGAAAAATATTAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTATATATGATACAAAAAAGATAGGTATATTCCAGATAAGACTTCTA													
CD111_S1_1	GATGCCTCATACACAGAAAAATATTAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTATATATGATACAAAAAAGATAGGTATATTCCAGATAAGACTTCTA													
Consensus	GATGCCTCATACACAGAAAAATATTAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTATATATGATACAAAAAAGATAGGTATATTCCAGATAAGACTTCTA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	TCGATGGTTGGAGGGGATTGAGGCTTGACTATGGTAATTATTATGCATCTAATCATTCTATGACCCTAGCAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCATGACGATGT													
CD111_S1_1	TCGATGGTTGGAGGGGATTGAGGCTTGACTATGGTAATTATTATGCATCTAATCATTCTATGACCCTAGCAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCATGACGATGT													
Consensus	TCGATGGTTGGAGGGGATTGAGGCTTGACTATGGTAATTATTATGCATCTAATCATTCTATGACCCTAGCAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCATGACGATGT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	CAAGAAGGATGGGCAGGAATTCAAACTATTCCCCGCAATTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCAATTAAGTAAT													
CD111_S1_1	CAAGAAGGATGGGCAGGAATTCAAACTATTCCCCGCAATTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCAATTAAGTAAT													
Consensus	CAAGAAGGATGGGCAGGAATTCAAACTATTCCCCGCAATTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCAATTAAGTAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_S1	CGCAAGTTAAGGAAGGAGATAAAATTGA													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	ATGGATTGTTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCCAGGGTTGCAATCTACAAGCA														
CD111_S2_1	ATGGATTGTTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCCAGGGTTGCAATCTACAAGCA														
Consensus	ATGGATTGTTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCCAGGGTTGCAATCTACAAGCA														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG														
CD111_S2_1	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG														
Consensus	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	ATCAGTTTGGGGAATATTGTTTGGGCCATTCAAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTCGACAATATGGTACATGGTCCGGGTCAGCCACA														
CD111_S2_1	ATCAGTTTGGGGAATATTGTTTGGGCCATTCAAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTCGACAATATGGTACATGGTCCGGGTCAGCCACA														
Consensus	ATCAGTTTGGGGAATATTGTTTGGGCCATTCAAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTCGACAATATGGTACATGGTCCGGGTCAGCCACA														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACCTATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA														
CD111_S2_1	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACCTATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA														
Consensus	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACCTATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCCAATTCCGTGACCCAACCCACATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC														
CD111_S2_1	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCCAATTCCGTGACCCAACCCACATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC														
Consensus	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCCAATTCCGTGACCCAACCCACATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	AATATTGTATAAAGTAGAGATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTCAAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA														
CD111_S2_1	AATATTGTATAAAGTAGAGATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTCAAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA														
Consensus	AATATTGTATAAAGTAGAGATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTCAAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAGGATAGGTATATTCCCGATAAGACTTCTA														
CD111_S2_1	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAGGATAGGTATATTCCCGATAAGACTTCTA														
Consensus	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAGGATAGGTATATTCCCGATAAGACTTCTA														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	TCGATGGTTTGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT														
CD111_S2_1	TCGATGGTTTGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT														
Consensus	TCGATGGTTTGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT														
CD111_S2_1	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT														
Consensus	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT														
CD111_S2_1	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT														
Consensus	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA														
CD111_S2_1	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA														
Consensus	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG														
CD111_S2_1	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG														
Consensus	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA														
CD111_S2_1	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA														
Consensus	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA														
	1691	1700	1710	1720	1730	1740	1750	1760	1770						
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD111_S2	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAAACCTAAGATGAATTGGTCATTTGGTCACTCTTCTTATTGA														
CD111_S2_1	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAAACCTAAGATGAATTGGTCATTTGGTCACTCTTCTTATTGA														
Consensus	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAAACCTAAGATGAATTGGTCATTTGGTCACTCTTCTTATTGA														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_T2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_1	ATGGATTGTTTAAAAAAGTCCTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTCTCCATAATTTTATCTTTCACACAAAGTTTTCAGGGTTGCAATCTACAGCA													
CD111_T2_2	ATGGATTGTTTAAAAAAGTCCTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTCTCCATAATTTTATCTTTCACACAAAGTTTTCAGGGTTGCAATCTACAGCA													
Consensus	ATGGATTGTTTAAAAAAGTCCTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTCTCCATAATTTTATCTTTCACACAAAGTTTTCAGGGTTGCAATCTACAGCA													
CD111_T2	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCACCTCCTAATAACTGGATCAATGATCCCAATGCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
Consensus	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCACCTCCTAATAACTGGATCAATGATCCCAATGCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
CD111_T2	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	ATCAGTTTGGGGAATATTGTTTGGGCCATTTCAGTTTCACCGACTTGATTAAATTGGATCCCACTTGAAGCCCGAATCTATCCATCCGAAGTATTCGACAAATATGGTACATGGTCCGGGTGAGCCACA													
Consensus	ATCAGTTTGGGGAATATTGTTTGGGCCATTTCAGTTTCACCGACTTGATTAAATTGGATCCCACTTGAAGCCCGAATCTATCCATCCGAAGTATTCGACAAATATGGTACATGGTCCGGGTGAGCCACA													
CD111_T2	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	ATCTTACCCAAACAACAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAATACCCAGTCCAAACTATGCAATCCAGCCGACATATCCGATCCATTTCCTTCGTAATGGATCAAGCCCGATA													
Consensus	ATCTTACCCAAACAACAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAATACCCAGTCCAAACTATGCAATCCAGCCGACATATCCGATCCATTTCCTTCGTAATGGATCAAGCCCGATA													
CD111_T2	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCAAATTCGTCGACCCAAACACATGTTGGTTGGGTGAGGATGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAGGATTGGC													
Consensus	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCAAATTCGTCGACCCAAACACATGTTGGTTGGGTGAGGATGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAGGATTGGC													
CD111_T2	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	AATATTGTATAAAGTAGAGATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTATTGTCATGGTACAATGGATTA													
Consensus	AATATTGTATAAAGTAGAGATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTATTGTCATGGTACAATGGATTA													
CD111_T2	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	GATGCCTCATACAACAGAAAGATATTAAACATGCTCTTAAAGTTAGTCTTGATGTTACGAGGTTGAATACTATACCGTTGGTAATATGATACAAAAAAGATAGGTATATTCCCGATAAGACTTCTA													
Consensus	GATGCCTCATACAACAGAAAGATATTAAACATGCTCTTAAAGTTAGTCTTGATGTTACGAGGTTGAATACTATACCGTTGGTAATATGATACAAAAAAGATAGGTATATTCCCGATAAGACTTCTA													
CD111_T2	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAATCATTCTATGACCTTAGGAAGAAATCGAAGATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
Consensus	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAATCATTCTATGACCTTAGGAAGAAATCGAAGATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
CD111_T2	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	CAAGAAAGGATGGGCGGGAATTCAACTATTCCCCGCAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCATGGCTGTCGAGAAATTAGAACTCTAAGAGAGCAAAAGGTGCAATTAGTAAT													
Consensus	CAAGAAAGGATGGGCGGGAATTCAACTATTCCCCGCAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCATGGCTGTCGAGAAATTAGAACTCTAAGAGAGCAAAAGGTGCAATTAGTAAT													
CD111_T2	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAGTTTGGATAAGGCAGAGCCATTGATCCCAATTGGGCTAACCTTT													
Consensus	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAGTTTGGATAAGGCAGAGCCATTGATCCCAATTGGGCTAACCTTT													
CD111_T2	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD111_T2_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_T2_2	ATGCTCAGATGTATGCGCCATTAAAGGCTCAACGGTGCAGGTGGACTTGGGCCATTTGGGCTCTTAACCTTTGGCTTCTCAAATTTGGAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA													
Consensus	ATGCTCAGATGTATGCGCCATTAAAGGCTCAACGGTGCAGGTGGACTTGGGCCATTTGGGCTCTTAACCTTTGGCTTCTCAAATTTGGAGAATATACACCGGT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_S1	ATGGATTGTTTAAAAAAGTC	TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
CD111_T1	ATGGATTGTTTAAAAAAGTC	TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
CD111_S3	ATGGATTGTTTAAAAAAGTC	TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
CD111_D1	ATGGATTGTTTAAAAAAGTC	TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
CD111_S2	ATGGATTGTTTAAAAAAGTC	TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
CD111_T2	ATGGATTGTTTAAAAAAGTC	CTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
Consensus	ATGGATTGTTTAAAAAAGTC	C TTCTCTTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAAGTTTTTCCAGGGTGC	CAATCTACAGCA											
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_S1	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	AA	AA	AACTGGATCAATGATCCCAATG	CCCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
CD111_T1	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	CA	AA	AACTGGATCAATGATCCCAATG	CCCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
CD111_S3	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	AA	AA	AACTGGATCAATGATCCCAATG	CTCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
CD111_D1	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	AA	AA	AACTGGATCAATGATCCCAATG	CTCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
CD111_S2	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	AA	AA	AACTGGATCAATGATCCCAATG	CCCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
CD111_T2	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	AA	AA	AACTGGATCAATGATCCCAATG	CCCAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
Consensus	CGGTTGATGTGAAAAATGTT	CATAGAACTGGTTATCATTTTCAACCTCCT	a AA	AA	AACTGGATCAATGATCCCAATG	C CAATGTACTATATG	TGGTGCTATCATCTATTCTACCAATACAAATCCATATGG							
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_S1	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
CD111_T1	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
CD111_S3	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
CD111_D1	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
CD111_S2	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
CD111_T2	ATCAGT	TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	C	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
Consensus	ATCAGT	T TGGGGAAATATTGTTTGGGCCCATTCAGTTTCAAC	c	GACTTGATTAA	TGGATCCC	ACTTGAGGCCGGAAATCTATCCATCCGAGTATT	T	GACAAATATGGTACATGGTCCGGGT	CAGCCACA					
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_S1	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	AA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
CD111_T1	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	GA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
CD111_S3	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	GA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
CD111_D1	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	GA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
CD111_S2	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	GA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
CD111_T2	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	GA	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
Consensus	ATCTTACCCARACACAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAA	AACTATGCAATCCAGCC	g	CAATATCCGATCCATT	TCTTCGTAATGGATC	ARGCCCGATA								
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD111_S1	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
CD111_T1	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
CD111_S3	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
CD111_D1	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
CD111_S2	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
CD111_T2	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	AAAA	C	ARGGATTGGC				
Consensus	ACA	ACCCATTAA	TGTCGCTGACGTTAGCATTAA	CAGACCCAA	TCCGTGACCCAA	ACCACATGTTGGTTGGGT	CAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGG	g AAAA	c	ARGGATTGGC				
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD111_S1	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
CD111_T1	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
CD111_S3	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
CD111_D1	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
CD111_S2	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
CD111_T2	AA	TATTGTATAAAAGTA	AA	ATTT	CATGAATGGACCAAG	ATT	CA	CA	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	CCAGTGT	TATTGCATGGTACAAATGGATTG
Consensus	AA	TATTGTATAAAAGTA	a A	ATTT	c ATGAATGGACCAAG	T C	AA	c A	ATCCACTTCATT	CAGTTGACGGTACTGGA	AA	TGGGAATGTCCTGATTTTTT	t CCAGTGT	TATTGCATGGTACAAATGGATT g
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD111_S1	GATGCC	TCATACACAGAA	AA	TATTAACATG	CTCTTAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	AGTTGGT	ATATATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA				
CD111_T1	GATGCC	TCATACACAGAA	AA	TATTAACATG	TTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	CGTTGGT	CA	TATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA			
CD111_S3	GATGCC	TCATACACAGAA	AA	TATTAACATG	TTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	AGTTGGT	AT	ATATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA			
CD111_D1	GATGCC	TCATACACAGAA	AA	TATTAACATG	TTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	AGTTGGT	AT	ATATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA			
CD111_S2	GATGCC	TCATACACAGAA	AA	TATTAACATG	CTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	CGTTGGT	AA	ATATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA			
CD111_T2	GATGCC	TCATACACAGAA	GA	TATTAACATG	CTCTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	CGTTGGT	AA	ATATGATACAAAAA	AGATAGGTATATTCC	AGATAGACTTCTA			
Consensus	GATGCC	TCATACACAGAA	a	TATTAACATG	c CTTAAAGTTAGTCTTGATGTTACGAGGTTTGA	ATACTATAC	c GTTGGT	aa	ATATGATACAAAAA	a AGATAGGTATATTCC	a	AGATAGACTTCTA		
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD111_S1	TCGATGGTT	GGAAGGGATTGAGG	CTTGAC	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT								
CD111_T1	TCGATGGTT	GGAAGGGATTGAG	ACTTGA	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT								
CD111_S3	TCGATGGTT	GGAAGGGATTGAG	ACTTGAT	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATAT								
CD111_D1	TCGATGGTT	GGAAGGGATTGAG	ACTTGAT	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATAT								
CD111_S2	TCGATGGTT	GGAAGGGATTGAG	ACTTGA	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT								
CD111_T2	TCGATGGTT	GGAAGGGATTGAG	ACTTGA	TATGGTAATTATTATGCATCTAAATCATTCTA	TGACCTAG	CAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT								
Consensus	TCGATGGTT	g GAAGGGATTGAG	a CTTGA	TATGGTAATTATTATGCATCTAAATCATTCTA	t GACCTAG	c AAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGAT g T								
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD111_S1	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	GGTCC	AATGGCCTGT	CGAAGATTAGAAACT	CTA	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
CD111_T1	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	GGTCC	AATGGCCTGT	CGAAGATTAGAAACT	CTA	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
CD111_S3	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	AGTCC	AATGGCCTGT	CGAAGATTAGAAACT	TTT	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
CD111_D1	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	AGTCC	AATGGCCTGT	CGAAGATTAGAAACT	TTT	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
CD111_S2	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	GGTCC	AATGGCCTGT	CGAAGATTAGAAACT	CTA	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
CD111_T2	CAAGAA	GGGATGGGCA	GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	CAATT	GGTCC	AATGGCCTGT	CGAAGATTAGAAACT	CTA	AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
Consensus	CAAGAA	GGGATGGG C	g GG	AATTCAAACTATTCCCCGCAAA	TATGGCTTGATCCTAGTGGAAAG	g CAATT	g TCC	AATGGCCTGT	CGAAGATTAGAAACT	c T	a AGAGAGCAAAAGGTGCAATT	TAAGTAAT		
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD111_S1	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	CA	TTTGATCCCA	ATTGGG	ATAACCTTT		
CD111_T1	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	AC	TTTGATCCCA	ATTGGG	ATAACCTTT		
CD111_S3	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	CA	TTTGATCCCA	ATTGGG	ATAACCTTT		
CD111_D1	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	CA	TTTGATCCCA	ATTGGG	ATAACCTTT		
CD111_S2	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	CA	TTTGATCCCA	ATTGGG	ATAACCTTT		
CD111_T2	CGC	AGGTTAAAC	AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	TT	CTATTT	CAAGTTTGGAT	ATAGGCAGAGC	CA	TTTGATCCCA	ATTGGG	ATAACCTTT		
Consensus	CGC	AGGTTAAAC	c AAAGGAGATAAAATTG	AGGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATT	c T	CTATTT	t C	AGTTTGGAT	ATAGGCAGAG c	CA	TTTGATCCCA	a TTGGG c	ATAACCTTT	
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD111_S1	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAC	GGTGCAAGG	TGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				
CD111_T1	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAT	TGGTGCAAGG	GGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				
CD111_S3	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAC	GGTGCAAGG	TGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				
CD111_D1	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAC	GGTGCAAGG	TGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				
CD111_S2	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAC	GGTGCAAGG	TGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				
CD111_T2	ATGCTC	AGAGTGATGCGCCATTAA	AGGCTCAAC	GGTGCAAGG	TGGACTTGGGCCATT	TGGGCTCTT	AACTTTGGAGAGATATAC	GCCGGTATTTTTCAGAGT	TTTTAAG	GCCCA				

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_P541	MDCLKKSSFLSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNNWINDPNAPHYNGVYHLFYQNPYGSVMGNIYWAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_P541_1	MDCLKKSSFLSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNNWINDPNAPHYNGVYHLFYQNPYGSVMGNIYWAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
CD111_P542	MDCLKKSSFLSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNNWINDPNAPHYNGVYHLFYQNPYGSVMGNIYWAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
Consensus	MDCLKKSSFLSLPIFLLYFSIILSFNNGYNASHKVFPGLQSTSTVDYKNVHRTGYHFQPPNNWINDPNAPHYNGVYHLFYQNPYGSVMGNIYWAHSYSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_P541	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLQGDGYWRTLIGSV ^W EKKGLAILYKSRDFHKWTKVQDPLHSVDGTGNWECPDFFPYLLHGTNGL													
CD111_P541_1	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLQGDGYWRTLIGSV ^W EKKGLAILYKSRDFHKWTKVQDPLHSVDGTGNWECPDFFPYLLHGTNGL													
CD111_P542	ILLNNTPIILHTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLQGDGYWRTLIGSV ^C EKKGLAILYKSRDFHKWTKVQDPLHSVDGTGNWECPDFFPYLLHGTNGL													
Consensus	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLQGDGYWRTLIGSV ^W EKKGLAILYKSRDFHKWTKVQDPLHSVDGTGNWECPDFFPYLLHGTNGL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_P541	DASYNKKNIKH ^A LKVSLDVTRFEYTYVGKYDTKKDRYIPDKTSIDG ^W NGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETLEQKVQLSN													
CD111_P541_1	DASYNKKNIKH ^A LKVSLDVTRFEYTYVGKYDTKKDRYIPDKTSIDG ^W NGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETLEQKVQLSN													
CD111_P542	DASYNKKNIKH ^V LKVSLDVTRFEYTYVGKYDTKKDRYIPDKTSIDG ^K NGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETLEQKVQLSN													
Consensus	DASYNKKNIKH ^A LKVSLDVTRFEYTYVGKYDTKKDRYIPDKTSIDG ^W NGLRLDYGNYYASKSFYDLRKNRRIMHGWANESDTYNDDYKKGWAGIQTIPRKLWLDPSGKQLVQWPVEELETLEQKVQLSN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_P541	RKLNKGDKIEYKGITPAQADVEYTFSSSLDKAEPFDPNWANLYAQDYCAIKGSTVQGGGLGPFGLLTLAS ^Q NLEEYTPVFFRVFKAQNKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDVDTLNKTLSLR													
CD111_P541_1	RKLNKGDKIEYKGITPAQADVEYTFSSSLDKAEPFDPNWANLYAQDYCAIKGSTVQGGGLGPFGLLTLAS ^Q NLEEYTPVFFRVFKAQNKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDVDTLNKTLSLR													
CD111_P542	RKLNKGDKIEYKGITPAQADVEYTFSSSLDKAEPFDPNWANLYAQDYCAIKGSTVQGGGLGPFGLLTLAS ^R NLEEYTPVFFRVFKAQDKYKVLHCSDASRSTLKNDKTHYKPSFAGYVDVDTLNKTLSLR													
Consensus	RKLNKGDKIEYKGITPAQADVEYTFSSSLDKAEPFDPNWANLYAQDYCAIKGSTVQGGGLGPFGLLTLAS ^Q NLEEYTPVFFRVFKAQ#KYKVLHCSDASRSTLKNDKTHYKPSFAGYVDVDTLNKTLSLR													
	521	530	540	550	560	570	580	589						
CD111_P541	SLIDHSVYESFGAGGKTCITSRYVPTLAIYDNAHLFYFNNGTETIKIKSLNANTHGKPKMNWSFGHSS ^Y													
CD111_P541_1	SLIDHSVYESFGAGGKTCITSRYVPTLAIYDNAHLFYFNNGTETIKIKSLNANTHGKPKMNWSFGHSSI													
CD111_P542	SLIDHSVYESFGAGGKTCITSRYVPTLAIYDNAHLFYFNNGTETIKIKSLNANTHGKPKMNWSFGHSS ^Y													
Consensus	SLIDHSVYESFGAGGKTCITSRYVPTLAIYDNAHLFYFNNGTETIKIKSLNANTHGKPKMNWSFGHSS ^Y													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
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CD111_P541	ATGGATTGTTTAAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCC													
CD111_P541_1	ATGGATTGTTTAAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCC													
Consensus	ATGGATTGTTTAAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
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CD111_P541	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
CD111_P541_1	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
Consensus	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTCAACCTCCTAATAACTGGATCAATGATCCCAATGCCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
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CD111_P541	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAGTATTTCGACAATATGGTACATGGTCCGGGTCAGCCACA													
CD111_P541_1	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAGTATTTCGACAATATGGTACATGGTCCGGGTCAGCCACA													
Consensus	ATCAGTTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACCGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAGTATTTCGACAATATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
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CD111_P541	ATCTTACCCAACAACAAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACATATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
CD111_P541_1	ATCTTACCCAACAACAAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACATATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
Consensus	ATCTTACCCAACAACAAGGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCCAAGTCCAAAACATATGCAATCCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
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CD111_P541	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCAAATCCGCGACCCAAACCACATGTTGGTTGGGTGAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC													
CD111_P541_1	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCAAATCCGCGACCCAAACCACATGTTGGTTGGGTGAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC													
Consensus	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAAGACCCAAATCCGCGACCCAAACCACATGTTGGTTGGGTGAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAAAAAAGGATTGGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
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CD111_P541	AATATTGTATAAAGTAGAGATTTTATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA													
CD111_P541_1	AATATTGTATAAAGTAGAGATTTTATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA													
Consensus	AATATTGTATAAAGTAGAGATTTTATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTA													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
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CD111_P541	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAAGATAGGTATATTCCCGATAAGACTTCTA													
CD111_P541_1	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAAGATAGGTATATTCCCGATAAGACTTCTA													
Consensus	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACCGTTGGTAATATGATACAAAAAAGATAGGTATATTCCCGATAAGACTTCTA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
CD111_P541_1	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
Consensus	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAAATCATTCTATGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT													
CD111_P541_1	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT													
Consensus	CAAGAAAGGATGGGCGGGAATTCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTGGTCCAATGGCCTGTCGAAGAATTAGAAGCTCTAAGAGAGCAAAAGGTGCATTAAAGTAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT													
CD111_P541_1	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT													
Consensus	CGCAGTTAAACAAGGAGATAAATTGAAGTTAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTCAAGTTTGGATAAGGCAGAGCCATTTGATCCCAATTGGGCTAACCTTT													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA													
CD111_P541_1	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA													
Consensus	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAAGGTGGACTTGGGCCATTTGGGCTCTTAACTTTGGCTTCTCAAATTTGGAGAATATACACCGGTATTTTTCAGAGTTTTTAAGGCCCA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAAGCCATCATTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG													
CD111_P541_1	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAAGCCATCATTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG													
Consensus	AAATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAAGACAATGTACAAGCCATCATTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGTCTTTAAGG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
CD111_P541_1	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
Consensus	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
	1691	1700	1710	1720	1730	1740	1750	1760	1770					
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P541	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCACTCTTCTTATTGA													
CD111_P541_1	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCACTCTTCAATCACT													
Consensus	CTATCAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCACTCTTCAaacaca													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	ATGGATTGTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGTTGCAATCTACAAGCA													
CD111_P54_2	ATGGATTGTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGTTGCAATCTACAAGCA													
Consensus	ATGGATTGTTAAAAAGTCTTCTCTTTTTCTTTGCCAATTTTTTTGTTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGTTGCAATCTACAAGCA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAAAACTGGGATCAATGATCCCAATGCTCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
CD111_P54_2	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAACTAAGTGGATCAATGATCCCAATGCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
Consensus	CGGTTGATGTGAAAAATGTTTCATAGAACTGGTTATCATTTTTCAACCTCCTAACTAAGTGGATCAATGATCCCAATGCCCAATGTACTATAATGGTGTCTATCATCTATTCTACCAATACAATCCATATGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	ATCAGTATGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACGGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTTGACAAATATGGTACATGGTCCGGGTCAGCCACA													
CD111_P54_2	ATCAGTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACGGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTCGACAAATATGGTACATGGTCCGGGTCAGCCACA													
Consensus	ATCAGTTGGGGAAATATTGTTTGGGCCCATTCAGTTTCAACGGACTTGATTAAATTGGATCCCACTTGAGCCCGGAATCTATCCATCCGAAGTATTCGACAAATATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCAAGTCCAAACTATGCAATCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
CD111_P54_2	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCAAGTCCAAACTATGCAATCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
Consensus	ATCTTACCAACAACAAGCCATTATCCTCTACACCGGAATCGTCGATGCAAAAAATACCAAGTCCAAACTATGCAATCCAGCCGACATATCCGATCCATTTCTTCGTAATGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGTGACCCACACCATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAACAAAGGATTGGC													
CD111_P54_2	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGTGACCCACACCATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAACAAAGGATTGGC													
Consensus	ACAACCCATTAAATTGTCGCTGACGTTAGCATTAAACAGACCCAAATTCGTGACCCACACCATGTTGGTTGGGTCAGGATGGTTATTGGAGAACCTTGATAGGGAGTGTGTGGGAAACAAAGGATTGGC													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	AATATTGTATAAAGTAGAATTTTCATGAATGGACCAAGATTCAACATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
CD111_P54_2	AATATTGTATAAAGTAGAATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
Consensus	AATATTGTATAAAGTAGAATTTTCATGAATGGACCAAGGTCCAAGATCCACTTCATTTCAGTTGACGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTTATTGCATGGTACAATGGATTG													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	GATGCCTCATACAACAAGAAAAATATTAAACATGTTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTAATATGATACAAAAAAGATAGGTATATTCAGATAAGACTTCTA													
CD111_P54_2	GATGCCTCATACAACAAGAAAAATATTAAACATGCTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTAATATGATACAAAAAAGATAGGTATATTCAGATAAGACTTCTA													
Consensus	GATGCCTCATACAACAAGAAAAATATTAAACATGTTCTTAAGTTAGTCTTGATGTTACGAGGTTTGAATACTATACAGTTGGTAATATGATACAAAAAAGATAGGTATATTCAGATAAGACTTCTA													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	TCGATGGTTGGAAGGGATTGAGACTTGATTATGGTAATTATTATGCATCTAATCATTCTACGACCTAGCAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATAT													
CD111_P54_2	TCGATGGTTGGAACGGATTGAGACTTGACTATGGTAATTATTATGCATCTAATCATTCTAGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
Consensus	TCGATGGTTGGAAGGGATTGAGACTTGACTATGGTAATTATTATGCATCTAATCATTCTAGACCTTAGGAAAAATCGAAGAATTATGTGGGGTTGGGCTAATGAATCAGATACTGTCAATGACGATGT													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	CAAGAAAGGATGGGCGGGAAATCAAACTATTCCCCGCAAAATATGGCTTGATCCTAGTGGAAGCAATTAGTCCAATGGCTGTGGAAGAATTAGAAGCTTTAGAGAGCAAAAGGTGCAATTAAAGTAAT													
CD111_P54_2	CAAGAAAGGATGGGCGGGAAATCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTAGTCCAATGGCTGTGGAAGAATTAGAAGCTTAGAGAGCAAAAGGTGCAATTAAAGTAAT													
Consensus	CAAGAAAGGATGGGCGGGAAATCAAACTATTCCCCGCAAACTATGGCTTGATCCTAGTGGAAGCAATTAGTCCAATGGCTGTGGAAGAATTAGAAGCTTAGAGAGCAAAAGGTGCAATTAAAGTAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	CGCAGTTAAACAAGGAGATAAAATTGAAGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTTCAGTTTGGATAAGGCGAGGCCATTTGATCCCAATTGGGCTAACCTTT													
CD111_P54_2	CGCAGTTAAACAAGGAGATAAAATTGAAGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTTCAGTTTGGATAAGGCGAGGCCATTTGATCCCAATTGGGCTAACCTTT													
Consensus	CGCAGTTAAACAAGGAGATAAAATTGAAGTTAAAGGAATCACACCTGCACAGGCTGATGTTGAAGTGACATTCTCATTTTTCAGTTTGGATAAGGCGAGGCCATTTGATCCCAATTGGGCTAACCTTT													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAGGGTGGACTTGGGCCATTTGGGCTCTTAACCTTTGGCTTCTCAAAATTTGGAAGAATATACGCCGGTATTTTTTCAGAGTTTTTAAGACCCA													
CD111_P54_2	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAGGGTGGACTTGGGCCATTTGGGCTCTTAACCTTTGGCTTCTCAAAATTTGGAAGAATATACACCGGTATTTTTTCAGAGTTTTTAAGGCCCA													
Consensus	ATGCTCAAGATGTATGCGCCATTAAAGGCTCAACGGTGCAGGGTGGACTTGGGCCATTTGGGCTCTTAACCTTTGGCTTCTCAAAATTTGGAAGAATATACACCGGTATTTTTTCAGAGTTTTTAAGGCCCA													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	AGATAAATATAAGGTTCTCATGTGCTCTGATGCCAAGATCAACCTCAAGAATGATAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGCTTTTAAGG													
CD111_P54_2	AGATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGCTTTTAAGG													
Consensus	AGATAAATATAAGGTTCTCATGTGCTCTGATGCATCAAGATCAACCTCAAGAATGATAGACAATGTACAGCCATCATTTGCTGGATATGTAGATGTAGATTTAACAAACAAGACATTGCTTTTAAGG													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
CD111_P54_2	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
Consensus	AGTTTGATTGATCACTCAGTAGTAGAAGCTTTGGTGCCGGAGGAAAAACATGTATCACATCAAGGGTTTATCCGACATTAGCAATTTATGATAATGCACATCTATTTGTCTTTAATAATGGGACAGAGA													
	1691	1700	1710	1720	1730	1740	1750	1760	1770					
CD111_P40_1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P54_1	CTATCAAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCGCTCTTCTTATTGA													
CD111_P54_2	CTATCAAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCACCTCTTCTTATTGA													
Consensus	CTATCAAAATTAAGTCATTGAATGCATGGACCATGGGTAACCTAAGATGAATTGGTCATTTGGTCaCTCTTCTTATTGA													

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_S1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIIWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_S2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_S2_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_S2_2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_S3	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDASKVHRTGYHFQPSKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_S3_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDASKVHRTGYHFQPSKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIIWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_S1	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSYRNHRGKVIMYKSNKNFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKDG													
CD141_S2	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSYRNHRGKVIMYKSDKDFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_S2_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSYRNHRGKVIMYKSDKDFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_S2_2	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADRTINKSQFRDPTTAWHGRDGNWRILVGSYRNHRGKVIMYKSDKDFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_S3	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDRNWRILVGSYRNHRGKVIMYKSNKNFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKDG													
CD141_S3_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDRNWRILVGSYRNHRGKVIMYKSNKNFMKWTAKAHLHSAPGTGNWECPDFFPVSLKNKDG													
Consensus	ILPGNKPVILYTGIVDANKTQVQNYAIPANISDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSYRNHRGKVIMYKS#K#FMKWTAKAHLHSAPGTGNWECPDFFPVSLKNK#G													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_S1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKHFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTRDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_S2	LDSYNGKD IKHYLVKVSFDYTRFDHYTYGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_S2_1	LDSYNGKD IKHYLVKVSFDYTRFDHYTYGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_S2_2	LDSYNGKD IKHYLVKVSFDYTRFDHYTYGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_S3	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_S3_1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
Consensus	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTVDNDYRKGWAGVHPIPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_S1	NKKLNKGEKVEIKGITVAQADVEVIFS FASLEKAELFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_S2	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_S2_1	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_S2_2	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_S3	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_S3_1	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
Consensus	NKKLNKGEKVEIKGITVAQADVEVIFS FSLDKAE PFDP SWADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAQDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
	521	530	540	550	560	570	580	582						
CD141_S1	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_S2	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_S2_1	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_S2_2	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_S3	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_S3_1	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
Consensus	RSLIDHSYVESFGAGGKTCITSRYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_D1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_D1_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_D1_3	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_D1_2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_D2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_D1	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_D1_1	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_D1_3	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_D1_2	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_D2	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
Consensus	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_D1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_D1_1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_D1_3	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_D1_2	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_D2	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
Consensus	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESD TVDNDYRKGWAGVHP IPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_D1	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_D1_1	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_D1_3	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_D1_2	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_D2	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
Consensus	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
	521	530	540	550	560	570	580	582						
CD141_D1	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_D1_1	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_D1_3	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_D1_2	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_D2	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
Consensus	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_T1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_T1_2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_T2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_T2_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_T3	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_T1_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAIWGNIVWAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_T1	ILPGNKPVILYTGIYDANKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_T1_2	ILPGNKPVILYTGIYDANKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_T2	ILPGNKPVILYTGIYDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_T2_1	ILPGNKPVILYTGIYDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_T3	ILPGNKPVILYTGIYDGNKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
CD141_T1_1	ILPGNKPVILYTGIYDANKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
Consensus	ILPGNKPVILYTGIYDANKTQVQNYAIPANMSDPYLRKWKIPDNNPLIVADKTINKSQFRDPTTAHWGRDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHPHLSAPGTGNWECPDFFPVSLKNKDG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_T1	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
CD141_T1_2	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
CD141_T2	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
CD141_T2_1	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
CD141_T3	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
CD141_T1_1	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
Consensus	LDSYNGKDIAHYLVKVSFDYTRFDHYTIGTYDTKKDQYFPDSTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIHLDPGKQLVQWPVQELETLRKKKYQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_T1	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_T1_2	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_T2	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_T2_1	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_T3	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_T1_1	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
Consensus	NKKLNKGEKVEIKGITVAQADVEVIFSFTSLDKAEPFDPADLYAQDVCAIKGSTVQGGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
	521	530	540	550	560	570	580	582						
CD141_T1	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_T1_2	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_T2	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_T2_1	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_T3	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_T1_1	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
Consensus	RSLIDHSIYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCATAAAGTTTATATGCACTTGCAATCTACTACTAGCC													
CD141_S2_2	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCAAAAGTTTATATGCACTTGCAATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCAAAAGTTTATATGCACTTGCAATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACAACCCAAAGGG													
CD141_S2_2	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACAACCCAAAGGG													
Consensus	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACAACCCAAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
CD141_S2_2	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAGACACAAGTCCAAACTATGCAATCCCAGCTAACATATCTGATCCATATCTTCGTAAGTGGATCAAGCCCAGTA													
CD141_S2_2	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAGACACAAGTCCAAACTATGCAATCCCAGCTAACATATCTGATCCATATCTTCGTAAGTGGATCAAGCCCAGTA													
Consensus	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAGACACAAGTCCAAACTATGCAATCCCAGCTAACATATCTGATCCATATCTTCGTAAGTGGATCAAGCCCAGTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
CD141_S2_2	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	TATAATGTACAAAAGTGATAAGGACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCCGCCCGGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTCAATGAAAATAAAATGGT													
CD141_S2_2	TATAATGTACAAAAGTGATAAGGACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCCGCCCGGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTCAATGAAAATAAAATGGT													
Consensus	TATAATGTACAAAAGTGATAAGGACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCCGCCCGGGTACTGGAATTTGGGAATGTCCTGATTTTTTCCAGTGTCAATGAAAATAAAATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	TTGGACACGTCATACATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATAACACTT													
CD141_S2_2	TTGGACACGTCATACATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATAACACTT													
Consensus	TTGGACACGTCATACATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATAACACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCTTTGATAATGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
CD141_S2_2	CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCTTTGATAATGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
Consensus	CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCTTTGATAATGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD141_S2	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_S2_1	TGTGAGGAAGGATGGGCTGGAGTTCAACCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAACAATTTGGTTCAATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
CD141_S2_2	TGTGAGGAAGGATGGGCTGGAGTTCAACCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAACAATTTGGTTCAATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
Consensus	TGTGAGGAAGGATGGGCTGGAGTTCAACCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAACAATTTGGTTCAATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD141_S2	-----+													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_S3	-----													
CD141_S3_1	ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTTTTGCCAATTCCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACCTTGCATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTTTTGCCAATTCCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACCTTGCATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_S3	-----													
CD141_S3_1	ATGTTGATGCTAGCAAGGTCCATAGAAGTGGTTATCATTTCACCTTCTAAAACTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTATTCTACCAAGTACACCCCAAGGG													
Consensus	ATGTTGATGCTAGCAAGGTCCATAGAAGTGGTTATCATTTCACCTTCTAAAACTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTATTCTACCAAGTACACCCCAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_S3	-----													
CD141_S3_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_S3	-----													
CD141_S3_1	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAAGACACAAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCAGTA													
Consensus	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAAGACACAAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCAGTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD141_S3	-----													
CD141_S3_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATAGAAATTGGAGAACTCTGGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATAGAAATTGGAGAACTCTGGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD141_S3	-----													
CD141_S3_1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAAATAAGATGGT													
Consensus	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAAATAAGATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD141_S3	-----													
CD141_S3_1	TTGGACACGTCATACATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATAAATACTTTCCGGATAACACTT													
Consensus	TTGGACACGTCATACATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATAAATACTTTCCGGATAACACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD141_S3	-----													
CD141_S3_1	CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAAGACATTCTTTGATAGTGGAAGAAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGGTTGATAACGA													
Consensus	CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAAGACATTCTTTGATAGTGGAAGAAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGGTTGATAACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD141_S3	-----													
CD141_S3_1	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAACAATTTGGTTCAATGGCCTGTTCAAGAACTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
Consensus	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAACAATTTGGTTCAATGGCCTGTTCAAGAACTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD141_S3	-----													
CD141_S3_1	AACAAAAAGTTGAACAAGGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTCAACAGTTTGGATAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
Consensus	AACAAAAAGTTGAACAAGGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTCAACAGTTTGGATAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD141_S3	-----													
CD141_S3_1	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTTGGGCTCCTAACTTTGGCTTCTAAAACTTAGAAGAAATACACACCCGTTTCTTTAGAAATTTTCAAGGC													
Consensus	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTTGGGCTCCTAACTTTGGCTTCTAAAACTTAGAAGAAATACACACCCGTTTCTTTAGAAATTTTCAAGGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
CD141_S3	-----													
CD141_S3_1	TCAAGATAGATACAAGTTCTTATGTGTTCCGATGCTTCAAGGTCAAGCCTAAGAATGAACAACACTATGTACAACCATCATTTGCTGGATATGTAGATGTAGATTAGCGGACAGAAATTGTCTCTT													
Consensus	TCAAGATAGATACAAGTTCTTATGTGTTCCGATGCTTCAAGGTCAAGCCTAAGAATGAACAACACTATGTACAACCATCATTTGCTGGATATGTAGATGTAGATTAGCGGACAGAAATTGTCTCTT													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
CD141_S3	-----													
CD141_S3_1	AGAAGTTTGATTGATCATTTCGGTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGGGTTTATCCACGTTGGCGATATTTGACAAGGCACATTTATTTGCATTCAATAACGGCGCGG													
Consensus	AGAAGTTTGATTGATCATTTCGGTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGGGTTTATCCACGTTGGCGATATTTGACAAGGCACATTTATTTGCATTCAATAACGGCGCGG													
	1691	1700	1710	1720	1730	1740	1749							
CD141_S3	-----													
CD141_S3_1	AGAGAATCACAATTGAGACTCTAATGCTTGAGCATGGCTAATGCAAGTTGCACTAG													
Consensus	AGAGAATCACAATTGAGACTCTAATGCTTGAGCATGGCTAATGCAAGTTGCACTAG													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130	
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CD141_D1	ATGGAGATTTTAAGAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC														
CD141_D1_2	ATGGAGATTTTAAGAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC														
CD141_D1_3	ATGGAGATTTTAAGAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC														
CD141_D1_1	ATGGAGATTTTAAGAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC														
Consensus	ATGGAGATTTTAAGAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC														
	131	140	150	160	170	180	190	200	210	220	230	240	250	260	
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CD141_D1	ATGTTGATGCTAGCAAGGTCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAAATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGG														
CD141_D1_2	ATGTTGATGCTAGCAAGGTCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAAATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGG														
CD141_D1_3	ATGTTGATGCTAGCAAGGTCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAAATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGG														
CD141_D1_1	ATGTTGATGCTAGCAAGGTCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAAATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGG														
Consensus	ATGTTGATGCTAGCAAGGTCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAAATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGG														
	261	270	280	290	300	310	320	330	340	350	360	370	380	390	
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CD141_D1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAAGTATGGTACATGGTCTGGGTCAAGCCACA														
CD141_D1_2	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAAGTATGGTACATGGTCTGGGTCAAGCCACA														
CD141_D1_3	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAAGTATGGTACATGGTCTGGGTCAAGCCACA														
CD141_D1_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAAGTATGGTACATGGTCTGGGTCAAGCCACA														
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAAGTATGGTACATGGTCTGGGTCAAGCCACA														
	391	400	410	420	430	440	450	460	470	480	490	500	510	520	
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CD141_D1	ATCTTGCCAGGCACAAAGCCTGTTATTCTCTACACTGGAAATGTGGATGGTAATAGACACAAAGTCCAAGTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCGATA														
CD141_D1_2	ATCTTGCCAGGCACAAAGCCTGTTATTCTCTACACTGGAAATGTGGATGGTAATAGACACAAAGTCCAAGTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCGATA														
CD141_D1_3	ATCTTGCCAGGCACAAAGCCTGTTATTCTCTACACTGGAAATGTGGATGGTAATAGACACAAAGTCCAAGTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCGATA														
CD141_D1_1	ATCTTGCCAGGCACAAAGCCTGTTATTCTCTACACTGGAAATGTGGATGGTAATAGACACAAAGTCCAAGTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCGATA														
Consensus	ATCTTGCCAGGCACAAAGCCTGTTATTCTCTACACTGGAAATGTGGATGGTAATAGACACAAAGTCCAAGTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCGATA														
	521	530	540	550	560	570	580	590	600	610	620	630	640	650	
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CD141_D1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGGCCAATTTCTGTATCCAAACACCGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT														
CD141_D1_2	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGGCCAATTTCTGTATCCAAACACCGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT														
CD141_D1_3	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGGCCAATTTCTGTATCCAAACACCGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT														
CD141_D1_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGGCCAATTTCTGTATCCAAACACCGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT														
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGGCCAATTTCTGTATCCAAACACCGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT														
	651	660	670	680	690	700	710	720	730	740	750	760	770	780	
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CD141_D1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTAAGTGGGAATTTGGGAATGTCTTGATTTTTTTCCAGTGTCTTAAAAATAAAGATGGT														
CD141_D1_2	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTAAGTGGGAATTTGGGAATGTCTTGATTTTTTTCCAGTGTCTTAAAAATAAAGATGGT														
CD141_D1_3	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTAAGTGGGAATTTGGGAATGTCTTGATTTTTTTCCAGTGTCTTAAAAATAAAGATGGT														
CD141_D1_1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTAAGTGGGAATTTGGGAATGTCTTGATTTTTTTCCAGTGTCTTAAAAATAAAGATGGT														
Consensus	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTAAGTGGGAATTTGGGAATGTCTTGATTTTTTTCCAGTGTCTTAAAAATAAAGATGGT														
	781	790	800	810	820	830	840	850	860	870	880	890	900	910	
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CD141_D1	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT														
CD141_D1_2	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT														
CD141_D1_3	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT														
CD141_D1_1	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT														
Consensus	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT														
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
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CD141_D1	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACTATTACGCGTCCAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
CD141_D1_2	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACTATTACGCGTCCAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
CD141_D1_3	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACTATTACGCGTCCAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
CD141_D1_1	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACTATTACGCGTCCAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
Consensus	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACTATTACGCGTCCAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
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CD141_D1	TGTGAGGAAGGATGGGCCGGAGTTCACCCATTCTCTGTAATAATTTGGCTTGATCCTAGTGGAAACAAATTGGTTCATGGCCTGTTCAAGAATTAGAARCTTAGAAGAGAAAAGGTCCAATTAAAT														
CD141_D1_2	TGTGAGGAAGGATGGGCCGGAGTTCACCCATTCTCTGTAATAATTTGGCTTGATCCTAGTGGAAACAAATTGGTTCATGGCCTGTTCAAGAATTAGAARCTTAGAAGAGAAAAGGTCCAATTAAAT														
CD141_D1_3	TGTGAGGAAGGATGGGCCGGAGTTCACCCATTCTCTGTAATAATTTGGCTTGATCCTAGTGGAAACAAATTGGTTCATGGCCTGTTCAAGAATTAGAARCTTAGAAGAGAAAAGGTCCAATTAAAT														
CD141_D1_1	TGTGAGGAAGGATGGGCCGGAGTTCACCCATTCTCTGTAATAATTTGGCTTGATCCTAGTGGAAACAAATTGGTTCATGGCCTGTTCAAGAATTAGAARCTTAGAAGAGAAAAGGTCCAATTAAAT														
Consensus	TGTGAGGAAGGATGGGCCGGAGTTCACCCATTCTCTGTAATAATTTGGCTTGATCCTAGTGGAAACAAATTGGTTCATGGCCTGTTCAAGAATTAGAARCTTAGAAGAGAAAAGGTCCAATTAAAT														
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
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CD141_D1	AACAAAAGTTGAACAGGGAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTACAAAGTTTGGACAGGGCAGAGCCATTTGACCTAGTTGGGCTGATC														
CD141_D1_2	AACAAAAGTTGAACAGGGAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTACAAAGTTTGGACAGGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
CD141_D1_3	AACAAAAGTTGAACAGGGAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTACAAAGTTTGGACAGGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
CD141_D1_1	AACAAAAGTTGAACAGGGAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTACAAAGTTTGGACAGGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
Consensus	AACAAAAGTTGAACAGGGAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTCATTACAAAGTTTGGACAGGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
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CD141_D1	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTACTAACTTTGGCTTCTAAAACTTAGAGAATACACACCCGTTTCTTCAGAAATTTTCAGGC														
CD141_D1_2	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTACTAACTTTGGCTTCTAAAACTTAGAGAATACACACCCGTTTCTTCAGAAATTTTCAGGC														
CD141_D1_3	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTACTAACTTTGGCTTCTAAAACTTAGAGAATACACACCCGTTTCTTCAGAAATTTTCAGGC														
CD141_D1_1	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTACTAACTTTGGCTTCTAAAACTTAGAGAATACACACCCGTTTCTTCAGAAATTTTCAGGC														
Consensus	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTACTAACTTTGGCTTCTAAAACTTAGAGAATACACACCCGTTTCTTCAGAAATTTTCAGGC														
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
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CD141_D1	TCATGATAAATACAAAGTTCCTATGTGTTCCGATGCCTCAAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAATTGCTCTC														
CD141_D1_2	TCATGATAAATACAAAGTTCCTATGTGTTCCGATGCCTCAAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAATTGCTCTC														
CD141_D1_3	TCATGATAAATACAAAGTTCCTATGTGTTCCGATGCCTCAAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAATTGCTCTC														
CD141_D1_1	TCATGATAAATACAAAGTTCCTATGTGTTCCGATGCCTCAAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAATTGCTCTC														
Consensus	TCATGATAAATACAAAGTTCCTATGTGTTCCGATGCCTCAAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAATTGCTCTC														
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
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CD141_D1	AGAAGTTTGATTGATCATTGATAGTGGAAAGTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGAGTTTATCCACAGTTGGCGATATTTGACAAGGCACATTTATTTGCATTACACACGGCGCGG														
CD141_D1_2	AGAAGTTTGATTGATCATTGATAGTGGAAAGTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGAGTTTATCCACAGTTGGCGATATTTGACAAGGCACATTTATTTGCATTACACACGGCGCGG														
CD141_D1_3	AGAAGTTTGATTGATCATTGATAGTGGAAAGTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGAGTTTATCCACAGTTGGCGATATTTGACAAGGCACATTTATTTGCATTACACACGGCGCGG														
CD141_D1_1	AGAAGTTTGATTGATCATTGATAGTGGAAAGTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGAGTTTATCCACAGTTGGCGATATTTGACAAGGCACATTTATTTGCATTACACACGGCGCGG														
Consensus	AGAAGTTTGATTGATCATTGATAGTGGAAAGTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGAGTTTATCCACAGTTGGCGATATTTGACAAGGCACATTTATTTGCATTACACACGGCGCGG														
	1691	1700	1710	1720	1730	1740	1749								
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CD141_D1	AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTTGCACTAG														
CD141_D1_2	AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTTGCACTAG														
CD141_D1_3	AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTTGCACTAG														
CD141_D1_1	AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTTGCACTAG														
Consensus	AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTTGCACTAG														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
CD141_T1_1	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGGGTGTATCATTTATTCTACCAAGTACAACCCAAGGG													
CD141_T1_1	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACAACCCAAGGG													
Consensus	ATGTTGATGTTAGCAAGGTCATAGAAGTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACAACCCAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
CD141_T1_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACCCGGAATTGTGGATGCTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
CD141_T1_1	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACCCGGAATTGTGGATGCTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
Consensus	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACCCGGAATTGTGGATGCTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
CD141_T1_1	ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAGCCAAATTCGTGATCCAACAACAGCATGGATGGGCCGAGATGGAATTTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	TATAATGTACAAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTACTGGAATTTGGGATGTCTGATTTTTTTCCAGTGTCAATAAAAATAAGATGGT													
CD141_T1_1	TATAATGTACAAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTACTGGAATTTGGGATGTCTGATTTTTTTCCAGTGTCAATAAAAATAAGATGGT													
Consensus	TATAATGTACAAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCCGGTACTGGAATTTGGGATGTCTGATTTTTTTCCAGTGTCAATAAAAATAAGATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAGGATCAGTACTTTCCGGATAGCACTT													
CD141_T1_1	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAGGATCAGTACTTTCCGGATAGCACTT													
Consensus	TTGGACACGTCATACACGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAGGATCAGTACTTTCCGGATAGCACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	CTATTGATGGATGGAAGGATTGAGACTCGACTATGGTAATATTACGCGTCCAGACATTCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
CD141_T1_1	CTATTGATGGATGGAAGGATTGAGACTCGACTATGGTAATATTACGCGTCCAGACATTCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
Consensus	CTATTGATGGATGGAAGGATTGAGACTCGACTATGGTAATATTACGCGTCCAGACATTCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD141_T1	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T1_2	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAARAACATTGGTTCATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
CD141_T1_1	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAARAACATTGGTTCATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
Consensus	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAARAACATTGGTTCATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAGGTCCAATTAAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD141_T1														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	ATGGAGATTTTAAGAGATCTTCTTCTTTGGGTTTGGCAATTCTTTGTTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC													
CD141_T2_1	ATGGAGATTTTAAGAGATCTTCTTCTTTGGGTTTGGCAATTCTTTGTTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAGATCTTCTTCTTTGGGTTTGGCAATTCTTTGTTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	ATGTTGATGCTAGCAAGGTCCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCAGTACAACCCAAGGG													
CD141_T2_1	ATGTTGATGCTAGCAAGGTCCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCAGTACAACCCAAGGG													
Consensus	ATGTTGATGCTAGCAAGGTCCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCAGTACAACCCAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGACCCGCTATCTACCCGTCCAAGTATTTGACAGTATGGTACATGGTCTGGGTCAGCCACA													
CD141_T2_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGACCCGCTATCTACCCGTCCAAGTATTTGACAGTATGGTACATGGTCTGGGTCAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGACCCGCTATCTACCCGTCCAAGTATTTGACAGTATGGTACATGGTCTGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	ATCTTGCCAGGCCAACAGCCTGTTATTCTCTACACTGGAATTGTGGATGGTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
CD141_T2_1	ATCTTGCCAGGCCAACAGCCTGTTATTCTCTACACTGGAATTGTGGATGGTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
Consensus	ATCTTGCCAGGCCAACAGCCTGTTATTCTCTACACTGGAATTGTGGATGGTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTTCTGTATCCAACAACCGCATGGATGGCCGAGATGGAATTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
CD141_T2_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTTCTGTATCCAACAACCGCATGGATGGCCGAGATGGAATTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTTCTGTATCCAACAACCGCATGGATGGCCGAGATGGAATTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT													
CD141_T2_1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT													
Consensus	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	TTGGACACGTCATACACGGCAAGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT													
CD141_T2_1	TTGGACACGTCATACACGGCAAGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT													
Consensus	TTGGACACGTCATACACGGCAAGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACATTTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA													
CD141_T2_1	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACATTTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA													
Consensus	CTATTGATGGATGGAAAGGATTGAGACTCGACTATGGTAACATTTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAATAATTTGGCTTGATCCTAGTGGAACAATTGGTTCAATGGCTGTTCAAGAATTAGAARTCTAGAAGAGAAAAGGTCCAATTAAT													
CD141_T2_1	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAATAATTTGGCTTGATCCTAGTGGAACAATTGGTTCAATGGCTGTTCAAGAATTAGAARTCTAGAAGAGAAAAGGTCCAATTAAT													
Consensus	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAATAATTTGGCTTGATCCTAGTGGAACAATTGGTTCAATGGCTGTTCAAGAATTAGAARTCTAGAAGAGAAAAGGTCCAATTAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_T2	AACAAAAAGTTGAACAGGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGACAGGGCAGAGCCATTTGAT													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_T1	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	TCCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_T2	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_T3	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_D1	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_D2	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_S2	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_S3	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
CD141_S1	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
Consensus	ATGGAGATTTTAAAGAGATCTTCTCTCTTTGGGTTTGGCAATTCCTTTGTTGTG	CTCTTTATCACCAATGGAGTATTTGTTGATGCTTCTCA	CAAGGTTTATATGCACCTTGCATCTACTACTAGCC											
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_T1	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGTCCAATGTATTACATGGG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_T2	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGGCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_T3	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGCCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_D1	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGCCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_D2	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGCCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_S2	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGCCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_S3	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGTCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
CD141_S1	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGTCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
Consensus	ATGTTGATGTAGCAAGGTCATAGAACTGGTTATCATTTTCACCTCTA	AAAAATGGATCAACGATCCAATGGCCCAATGTATTACATGGAG	TGTATCATTTATCTACCAAGTACACCCCAAGGG											
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_T1	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_T2	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_T3	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_D1	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_D2	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_S2	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_S3	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
CD141_S1	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACATGGTCC	GGGTACGCCAC								
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCATTCCGGTCTCA	AAGGACTTGATCAATTGGATCCCACTTGAAACCCGCTAT	CTACCCGTC	CAAGGATTTGAC	CAAGTATGGTACAT									

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P181	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_P181_2	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_P181_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_P182_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
CD141_P182	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLQSTTSHYDVSKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYVAHSVSKDLINWIPLEPAIYPSKVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_P181	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_P181_2	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_P181_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_P182_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
CD141_P182	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
Consensus	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIPDNPLIYADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKHPLHSAPGTGNWECPDFFPVSLKNKNG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_P181	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P181_2	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P181_1	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P182_1	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P182	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
Consensus	LDSYNGKD IKHYLVKVSFDVTRFDHYTVGT YDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDNGKNRRILLGWANESDTYDNDYRKGWAGVHPIPRKIHLDP SGKQLVQWPVQELET LRKKKYQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_P181	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
CD141_P181_2	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
CD141_P181_1	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
CD141_P182_1	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
CD141_P182	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
Consensus	NKKLNKG EKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDVCAIKGSTVQGG LGPFGLLTLASKNLE EYTPVFFRIFKTHDKYKVLHCS DASRSSLKNETTHYKPSFAGYVDV DLADKKLSL													
	521	530	540	550	560	570	580	588	582					
CD141_P181	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_P181_2	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_P181_1	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_P182_1	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
CD141_P182	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													
Consensus	RSLIDHSVYESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITTIETLNAWSMANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P401	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSYSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
CD141_P401_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSYSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHKVYMHQLSTTSHVDASKVHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSYSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_P401	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIYADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIHYKSNKNFMKWTAKAKHPLHSAPGTGNWECPDFFPYSLKNKDG													
CD141_P401_1	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIYADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIHYKSNKNFMKWTAKAKHPLHSAPGTGNWECPDFFPYSLKNKDG													
Consensus	ILPGNKPVILYTGIVDGNKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIYADKTINKSQFRDPTTAWHGRDGNWRILVGSVRNHRGKVIHYKSNKNFMKWTAKAKHPLHSAPGTGNWECPDFFPYSLKNKDG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_P401	LDSYNGKDIKHVLYKVSFDVTRFDHYTIGTYDTKKDQYFPDSTSIDGKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIWLDPGSKQLVQWPVQELETLRKKKVQLN													
CD141_P401_1	LDSYNGKDIKHVLYKVSFDVTRFDHYTIGTYDTKKDQYFPDSTSIDGKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIWLDPGSKQLVQWPVQELETLRKKKVQLN													
Consensus	LDSYNGKDIKHVLYKVSFDVTRFDHYTIGTYDTKKDQYFPDSTSIDGKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDVTYDNDYRKGWAGVHPIPRKIWLDPGSKQLVQWPVQELETLRKKKVQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_P401	NKKLNKGKVEIKGITVAQADVEVIFSFTSLDKAEPDPSWADLYAQDYCAIKGSTVQGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKYLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
CD141_P401_1	NKKLNKGKVEIKGITVAQADVEVIFSFTSLDKAEPDPSWADLYAQDYCAIKGSTVQGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKYLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
Consensus	NKKLNKGKVEIKGITVAQADVEVIFSFTSLDKAEPDPSWADLYAQDYCAIKGSTVQGGLGPFGLLTLASKNLEEYTPVFFRIFKAHDKYKYLHCSDASRSSLKNETTHYKPSFAGYVDVLDADKKLSL													
	521	530	540	550	560	570	580	582						
CD141_P401	RSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITLNAWSHANAKLH													
CD141_P401_1	RSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITLNAWSHANAKLH													
Consensus	RSLIDHSIVESFGAGGKTCITSRVYPTLAIFDKAHLFAFNNGAERITITLNAWSHANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P541	MEILRRSSSLWVLPILLLCFFINNGVFVDASHK V YMHQLQSTTSHVDASKYHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
CD141_P541_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHK V YMHQLQSTTSHVDASKYHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
CD141_P542	MEILRRSSSLWVLPILLLCFFINNGVFVDASHK A YMHQLQSTTSHVDASKYHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
CD141_P542_1	MEILRRSSSLWVLPILLLCFFINNGVFVDASHK A YMHQLQSTTSHVDASKYHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
Consensus	MEILRRSSSLWVLPILLLCFFINNGVFVDASHK v YMHQLQSTTSHVDASKYHRTGYHFQPPKNWINDPNGPMYYNGVYHLFYQYNPKGAINGNIYWAHSVSKDLINWIPLEPAIYPSKYVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_P541	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHP LHSAPGTGNWECPDFFPV SL LKNKNG													
CD141_P541_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHP LHSAPGTGNWECPDFFPV SL LKNKNG													
CD141_P542	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHP LHSAPGTGNWECPDFFPV L LKNKNG													
CD141_P542_1	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHP LHSAPGTGNWECPDFFPV L LKNKNG													
Consensus	ILPGNKPVILYTGIVDANKTQVQNYAIPANMSDPYLRKWKIKPDNNPLIVADKTINKSQFRDPTTAHMG RDGNWRILVGSVRNHRGKVIMYKSNKNFMKWTAKAHP LHSAPGTGNWECPDFFPV s LKNKNG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_P541	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTYDND V RKGWAGVHP IPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P541_1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSIDGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTYDND V RKGWAGVHP IPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P542	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSYD GWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTYDNDARKGWAGVHP IPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
CD141_P542_1	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTSYD GWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTYDNDARKGWAGVHP IPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
Consensus	LDSYNGKD IKHYLVKVSFDYTRFDHYTIGTYDTKKDKYFPDNTS!DGWKGLRLDYGNYYASKTFFDSGKNRRILLGWANESDTYDND v RKGWAGVHP IPRKIWLDP SGKQLVQWPVQELET LRKKKYQLN													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_P541	NKKLNGGKKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDYCAIKGSTVQGG LGPFGLLTLASKNLEEYTPV F FRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_P541_1	NKKLNGGKKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDYCAIKGSTVQGG LGPFGLLTLASKNLEEYTPV F FRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_P542	NKKLNGGKKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDYCAIKGSTVQGG LGPFGLLTLASKNLEEYTPVSFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
CD141_P542_1	NKKLNGGKKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDYCAIKGSTVQGG LGPFGLLTLASKNLEEYTPVSFRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
Consensus	NKKLNGGKKVEIKGITVAQADVEVIFSFTSLDKAEPFDP SHADLYAQDYCAIKGSTVQGG LGPFGLLTLASKNLEEYTPV f FRIFKAHDKYKVLHCSDASRSSLKNETTHYKPSFAGYVDYDLADKKLSL													
	521	530	540	550	560	570	580	582						
CD141_P541	RSLIDHSVYESFGAGGK T CITSRVYPTLAIFDKAHL F VFNNGAERIT IETLNAWSMANAKLH													
CD141_P541_1	RSLIDHSVYESFGAGGK T CITSRVYPTLAIFDKAHL F VFNNGAERIT IETLNAWSMANAKLH													
CD141_P542	RSLIDHSVYESFGAGGK I CITSRVYPTLAIFDKAHL F AFNNGAERIT IETLNAWSMANAKLH													
CD141_P542_1	RSLIDHSVYESFGAGGK I CITSRVYPTLAIFDKAHL F AFNNGAERIT IETLNAWSMANAKLH													
Consensus	RSLIDHSVYESFGAGGK t CITSRVYPTLAIFDKAHL f VFNNGAERIT IETLNAWSMANAKLH													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P181	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181_2	ATGGAGATTTTAAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCCTTTGTTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
CD141_P181_1	ATGGAGATTTTAAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCCTTTGTTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAGATCTTCTCTCTTTGGGTTTTGCCAATTCCTTTGTTGTGTTTCCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAGTTTATATGCACTTGCAATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	ATGTTGATGTTAGCAGGTCCATAGAACTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCACTACAACCCAAAGGG													
CD141_P181_2	ATGTTGATGTTAGCAGGTCCATAGAACTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCACTACAACCCAAAGGG													
CD141_P181_1	ATGTTGATGTTAGCAGGTCCATAGAACTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCACTACAACCCAAAGGG													
Consensus	ATGTTGATGTTAGCAGGTCCATAGAACTGGTTATCATTTTCACCTCCTAAAAATTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTTATTCTACCACTACAACCCAAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	AGCGATATGGGGCAATATTGTTTGGGCCCATTGGTCTCAAGGACTTGATCAATTGGATCCCATTGAACCCGCTATCTACCCGTCCTAAAGTATTTGACAAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P181_2	AGCGATATGGGGCAATATTGTTTGGGCCCATTGGTCTCAAGGACTTGATCAATTGGATCCCATTGAACCCGCTATCTACCCGTCCTAAAGTATTTGACAAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P181_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTGGTCTCAAGGACTTGATCAATTGGATCCCATTGAACCCGCTATCTACCCGTCCTAAAGTATTTGACAAAGTATGGTACATGGTCCGGGTCAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTGGTCTCAAGGACTTGATCAATTGGATCCCATTGAACCCGCTATCTACCCGTCCTAAAGTATTTGACAAAGTATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	ATCTTGCCAGGCAACAGCCTGTTATTCTCTACACTGGAAATTGTGGATGCTAACAGACACAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAAGTGGATCAAGCCCGACA													
CD141_P181_2	ATCTTGCCAGGCAACAGCCTGTTATTCTCTACACTGGAAATTGTGGATGCTAACAGACACAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAAGTGGATCAAGCCCGACA													
CD141_P181_1	ATCTTGCCAGGCAACAGCCTGTTATTCTCTACACTGGAAATTGTGGATGCTAACAGACACAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAAGTGGATCAAGCCCGACA													
Consensus	ATCTTGCCAGGCAACAGCCTGTTATTCTCTACACTGGAAATTGTGGATGCTAACAGACACAGTCCAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAAGTGGATCAAGCCCGACA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACACAGCATGGATGGGCCGAGATGGAATTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
CD141_P181_2	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACACAGCATGGATGGGCCGAGATGGAATTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
CD141_P181_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACACAGCATGGATGGGCCGAGATGGAATTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCAACACAGCATGGATGGGCCGAGATGGAATTGGAGAATCTTGTTGGGAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCGCCCCGGGTACTGGAATTGGGAATGTCCTGATTTTTTCCAGTGTCAATTGAAAAATAAAATGGT													
CD141_P181_2	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCGCCCCGGGTACTGGAATTGGGAATGTCCTGATTTTTTCCAGTGTCAATTGAAAAATAAAATGGT													
CD141_P181_1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCGCCCCGGGTACTGGAATTGGGAATGTCCTGATTTTTTCCAGTGTCAATTGAAAAATAAAATGGT													
Consensus	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCGCCCCGGGTACTGGAATTGGGAATGTCCTGATTTTTTCCAGTGTCAATTGAAAAATAAAATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	TTGGACACGTCATACATGGCAAGACGTTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATACACTT													
CD141_P181_2	TTGGACACGTCATACATGGCAAGACGTTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATACACTT													
CD141_P181_1	TTGGACACGTCATACATGGCAAGACGTTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATACACTT													
Consensus	TTGGACACGTCATACATGGCAAGACGTTAAACATGTTCTTAAGGTTAGCTTTGATGTTACTAGGTTTGATCATTACACAGTTGGTACATATGACACCAAAAGGATAAATACTTTCCGGATACACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P181	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACTATTACGCGTCCAAGACATTCCTTGATAATGGCAGAAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA													
CD141_P181_2	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACTATTACGCGTCCAAGACATTCCTTGATAATGGCAGAAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA													
CD141_P181_1	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACTATTACGCGTCCAAGACAT													

		1	10	20	30	40	50	60	70	80	90	100	110	120	130	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTTTTGCCAATTCCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCAATCTACTACTAGCC														
CD141_P182_1		ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTTTTGCCAATTCCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCAATCTACTACTAGCC														
Consensus		ATGGAGATTTTAAGAAGATCTTCTTCTCTTTGGGTTTTGCCAATTCCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCAATCTACTACTAGCC														
		131	140	150	160	170	180	190	200	210	220	230	240	250	260	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		ATGTTGATGCTAGCAAGGTCCATAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGTCCAATGTATTACAATGGAGTGTACCATTTATTCTACCAGTACACCCAAAGGG														
CD141_P182_1		ATGTTGATGCTAGCAAGGTCCACAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAATGATCCAATGGCCAATGTATTACAATGGAGTGTATCATTTATTCTACCAGTACACCCAAAGGG														
Consensus		ATGTTGATGCTAGCAAGGTCCACAGAAGTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCAATGTATTACAATGGAGTGTACCATTTATTCTACCAGTACACCCAAAGGG														
		261	270	280	290	300	310	320	330	340	350	360	370	380	390	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		AGCGATATGGGGCAACATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA														
CD141_P182_1		AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA														
Consensus		AGCGATATGGGGCAACATTGTTTGGGCCCATTCGGTCTCAAGGACTTGATCAATTGGATCCCACCTTGAAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA														
		391	400	410	420	430	440	450	460	470	480	490	500	510	520	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAGACACAAGTCCAAAATTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA														
CD141_P182_1		ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAATAAGACACAAGTCCAAAATATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA														
Consensus		ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGCTAACAGACACAAGTCCAAAATATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA														
		521	530	540	550	560	570	580	590	600	610	620	630	640	650	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAAGCCAATTTTCGTGATCCACAAACCGCTTGGATGGGCCGAGATGGAATTGGAGAATCTTGGTAGGGAGTGTGAGGAATCATAGGGGAAGGT														
CD141_P182_1		ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAAGCCAATTTTCGTGATCCACAAACCGCTTGGATGGGCCGAGATGGAATTGGAGAATCTTGGTAGGGAGTGTGAGGAATCATAGGGGAAGGT														
Consensus		ACAATCCATTGATTGTTGCTGACAAAACCATCAACAAAAGCCAATTTTCGTGATCCACAAACCGCTTGGATGGGCCGAGATGGAATTGGAGAATCTTGGTAGGGAGTGTGAGGAATCATAGGGGAAGGT														
		651	660	670	680	690	700	710	720	730	740	750	760	770	780	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAATCCCACCTCCACTCGGCCCCGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT														
CD141_P182_1		TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAACACCCACTCCACTCAGCCCCGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTCAATGAAAAATAAATGGT														
Consensus		TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAAaACCCTCCACTCaGCCCCGGTACTGGAATTTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAaATGGT														
		781	790	800	810	820	830	840	850	860	870	880	890	900	910	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		TTGGACACGTCATACAAATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATAAGTACATTCCGGATAACACTT														
CD141_P182_1		TTGGACACGTCATACAAATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATAAATACATTCCGGATAACACTT														
Consensus		TTGGACACGTCATACAAATGGCAAGACATTAAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATAAaTACaATTCCGGATAACACTT														
		911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACCTATTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
CD141_P182_1		CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACCTATTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
Consensus		CTATTGATGGATGGAAGGATTGAGACTTGACTATGGTAACCTATTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACTGTTGATAACGA														
		1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		TGTGAGGAAGGATGGGCCGGAGTTCACCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAGAACATTGGTTCAATGGCCTGTTCAAGAATTAGAAGCTTAGAAGAGAAAAGGTCCAATTAAT														
CD141_P182_1		TGTGAGGAAGGATGGGCCGGAGTTCACCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAGAACATTGGTTCAATGGCCTGTTCAAGAATTAGAAGCTTAGAAGAGAAAAGGTCCAATTAAT														
Consensus		TGTGAGGAAGGATGGGCCGGAGTTCACCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAGAACATTGGTTCAATGGCCTGTTCAAGAATTAGAAGCTTAGAAGAGAAAAGGTCCAATTAAT														
		1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		AACAAAAAGTTGAACAAAGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCTCAGGCTGATGTTGAAGTGATTTTCTCATTCACAAGTTTGGATAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
CD141_P182_1		AACAAAAAGTTGAACAAAGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCAAGGCTGATGTTGAAGTGATTTTCTCATTCACAAGTTTGGATAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
Consensus		AACAAAAAGTTGAACAAAGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCaCAGGCTGATGTTGAAGTGATTTTCTCATTCACAAGTTTGGATAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC														
		1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		TTTACGCGCAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTCCTAACTTTGGCTTCTAAAAACGTAGAAGAATACACACCCGTTTTCTTTAGAATTTTCAGGGC														
CD141_P182_1		TTTACGCGCAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTCCTAACTTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTTAGAATTTTCAGGGC														
Consensus		TTTACGCGCAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTGGGCTCCTAACTTTGGCTTCTAAAAACgTAGAAGAATACACACCCGTTTTCTTTAGAATTTTCAGGGC														
		1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		TCAGATAAATACAAAGTTCTTATGTGTTCCGATGCTTCAGGTCAGCCTAAGAATGAACAACTATGTACAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAARTTGTCTCTT														
CD141_P182_1		TCATGATAAATACAAAGTTCTTATGTGTTCTGATGCTTCAGGTCAGCCTAAGAATGAACAACTATGTATAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAARTTGTCTCTT														
Consensus		TCAGATAAATACAAAGTTCTTATGTGTTCCGATGCTTCAGGTCAGCCTAAGAATGAACAACTATGTACaACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAARTTGTCTCTT														
		1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690	
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		AGAAGTTTGATTGATCATTTCGGTAGTGGAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAAGGGTTATCCAACGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACACACGGCGCGG														
CD141_P182_1		AGAAGTTTGATTGATCATTTCGGTAGTCGAAGTTTTGGTGCCGGAGGAAAAACATGCATTACATCAAGGGTTATCCAACATTGGCGATATTTGACAAGGCACATTTATTTGTGTTCAACACACGGCGCAG														
Consensus		AGAAGTTTGATTGATCATTTCGGTAGTcGAAGTTTTGGTGCCGGAGGAAAAACATGCATTACATCAAGGGTTATCCAACaTTGGCGATATTTGACAAGGCACATTTATTTGcaTTCAACACACGGCGCaG														
		1691	1700	1710	1720	1730	1740	1749								
		-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----														
CD141_P182		AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG														
CD141_P182_1		AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG														
Consensus		AGAGAATCACAAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG														

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCAATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAAGATCTTCTTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACTTGCAATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	ATGTTGATGCTAGCAAGGTCCATAGAACTGGTTATCATTTCACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTATTCTACCAGTACAACCCAAGGG													
Consensus	ATGTTGATGCTAGCAAGGTCCATAGAACTGGTTATCATTTCACCTCCTAAAACTGGATCAACGATCCAATGGCCCAATGTATTACAATGGAGTGTATCATTATTCTACCAGTACAACCCAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGACCCGCTATCTACCCGTCCAAAGTATTTGACAGTATGGTACATGGTCTGGGTCAAGCCACA													
Consensus	AGCGATATGGGGCAATATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACTTGACCCGCTATCTACCCGTCCAAAGTATTTGACAGTATGGTACATGGTCTGGGTCAAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGGTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
Consensus	ATCTTGCCAGGCAACAAGCCTGTTATTCTCTACACTGGAATTGTGGATGGTAATAAGACACAAGTCCAAACTATGCAATCCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTTTCGTGATCCAACAACCGCATGGATGGCCGAGATGGAATTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAATTTTCGTGATCCAACAACCGCATGGATGGCCGAGATGGAATTGGAGAATCTTGGTAGGCAGTGTGAGGAATCATAGGGGAAGGT													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT													
Consensus	TATAATGTACAAAGTAATAAGAACTTCATGAATGGACCAAGCCAACACCCACTCCACTCAGCCCCCGGTACTGGAATTGGGAATGTCCTGATTTTTTTCCAGTGTCAATAAAAATAAAGATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	TTGGACACGTCATACACGGCAAGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT													
Consensus	TTGGACACGTCATACACGGCAAGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAAGGTTTGATCATTACACAATTGGTACATATGACACCAAAAAGGATCAGTACTTTCCGGATAGCACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	CTATTGATGGATGGAAGGATTGAGACTCGACTATGGTAACATTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCATATGAATCAGATACTGTTGATAACGA													
Consensus	CTATTGATGGATGGAAGGATTGAGACTCGACTATGGTAACATTACGCGTCCAAGACATTCTTTGATAGTGGCAAGAATCGTAGGATTTTGTGGGTTGGGCATATGAATCAGATACTGTTGATAACGA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAATAATTTGGCTTGATCCTAGTGGAAAACAATTGGTTCAATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAAGGTCCAATTAAT													
Consensus	TGTGAGGAAGGATGGGCCGGAGTTCACCTATTCTCGTAATAATTTGGCTTGATCCTAGTGGAAAACAATTGGTTCAATGGCCTGTTCAAGAATTAGAAGTCTAAGAAGAAAAAGGTCCAATTAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	AACAAAAAGTTGAACAAGGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGACAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
Consensus	AACAAAAAGTTGAACAAGGGAGAAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGACAAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTTGGGCTACTAAGTTTGGCTTCTAAAACTTAGAAGAATACACACCCGTTTTCTTCAGAATTTTCAGGC													
Consensus	TTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAAGGTGGACTTGGGCCTTTTGGGCTACTAAGTTTGGCTTCTAAAACTTAGAAGAATACACACCCGTTTTCTTCAGAATTTTCAGGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	TCATGATAAATACAAAGTTCATTATGTGTTCCGATGCCCAAGGTCAGGCCTAAGAATGAACAACACTATGTATAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAGTTGTCTCTC													
Consensus	TCATGATAAATACAAAGTTCATTATGTGTTCCGATGCCCAAGGTCAGGCCTAAGAATGAACAACACTATGTATAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGGACAGAAGTTGTCTCTC													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	AGAAGTTTGATTGATCATTCGATAGTGGAAGTTTTGGTGCcGGAGGAAAAACATGCATTACATCAAGAGTTTATCCAACGTTGGCGATATTTGACAAAGCACATTTATTTGCATTCAACAACGGCGCGG													
Consensus	AGAAGTTTGATTGATCATTCGATAGTGGAAGTTTTGGTGCcGGAGGAAAAACATGCATTACATCAAGAGTTTATCCAACGTTGGCGATATTTGACAAAGCACATTTATTTGCATTCAACAACGGCGCGG													
	1691	1700	1710	1720	1730	1740	1749							
CD141_P401	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD141_P401_1	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
Consensus	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAa													

[illegible]

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
CD141_P542	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGCTTATATGCACCTTGCATCTACTACTAGCC													
CD141_P541	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTATATGCACCTTGCATCTACTACTAGCC													
CD141_P182	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTATATGCACCTTGCATCTACTACTAGCC													
CD141_P401	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTATATGCACCTTGCATCTACTACTAGCC													
CD141_P181	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTATATGCACCTTGCATCTACTACTAGCC													
Consensus	ATGGAGATTTTAAGAGATCTTCTTCTCTTTGGGTTTTGCCAATTCTTTTGTGTGCTTCTTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGCTTATATGCACCTTGCATCTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
CD141_P542	ATGTTGATGCAAGCAGGTCCTAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGTCCAATGTATTACAATGGAGTGTAACATTTATTCTACCAGTACAACCCAAGGG													
CD141_P541	ATGTTGATGCTAGCAGGTCCTACAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAATGATCCAATGGCCAATGTATTACAATGGAGTGTAACATTATTCTACCAGTACAACCCAAGGG													
CD141_P182	ATGTTGATGCTAGCAGGTCCTAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGTCCAATGTATTACAATGGAGTGTAACATTTATTCTACCAGTACAACCCAAGGG													
CD141_P401	ATGTTGATGCTAGCAGGTCCTAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCAATGTATTACAATGGAGTGTAACATTATTCTACCAGTACAACCCAAGGG													
CD141_P181	ATGTTGATGCTAGCAGGTCCTAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCAATGTATTACAATGGAGTGTAACATTATTCTACCAGTACAACCCAAGGG													
Consensus	ATGTTGATGCTAGCAGGTCCTAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATCAACGATCCAATGGCCAATGTATTACAATGGAGTGTAACATTATTCTACCAGTACAACCCAAGGG													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
CD141_P542	AGCGATATGGGGCAATTTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P541	AGCGATATGGGGCAATTTATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P182	AGCGATATGGGGCAATTTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P401	AGCGATATGGGGCAATTTATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
CD141_P181	AGCGATATGGGGCAATTTATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
Consensus	AGCGATATGGGGCAATTTATTGTTTGGGCCCATTCGGTCTCAAAGGACTTGATCAATTGGATCCCACCTTGAACCCGCTATCTACCCGTCCAAGTATTTGACAAGTATGGTACATGGTCCGGGTCAGCCACA													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
CD141_P542	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
CD141_P541	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
CD141_P182	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
CD141_P401	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
CD141_P181	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
Consensus	ATCTTGCCAGGCACACAGCCTGTTATTCTCTACACTGGAAATGTGGATGCTAATAGACACAAGTCCAAAATATGCAATCCCGGCTACATGTCTGATCCATATCTTCGTAAGTGGATCAAGCCCGATTA													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
CD141_P542	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACCGCTTGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
CD141_P541	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACCGCTTGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
CD141_P182	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACCGCTTGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
CD141_P401	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACCGATGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
CD141_P181	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACAGATGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
Consensus	ACAATCCATTGATTGTTGCTGACAAACCATCAACAAAGCCAAATTCGTGATCCACACACCGCTTGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGAGTGTGAGGAATCATAGGGGAAGGGA													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
CD141_P542	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAACCCACTCCACTCAGCTCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
CD141_P541	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAACCCACTCCACTCAGCCCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
CD141_P182	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAATCCCACTCCACTCAGCCCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
CD141_P401	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAACCCACTCCACTCAGCCCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
CD141_P181	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAACCCACTCCACTCAGCCCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
Consensus	TATAATGTACAAAGTAATAGAAGTTCATGAATGGACCAAGGCCAACAACCCACTCCACTCAGCCCCGGGTACTGGAAATTTGGGAATGTCTGATTTTTCAGTGTCTTGAATAAATATGGT													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
CD141_P542	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAATTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
CD141_P541	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAATTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
CD141_P182	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAATTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
CD141_P401	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAATTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
CD141_P181	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAGTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
Consensus	TTGGACACGTCATACAAATGGCAAGGACATTAACATGTTCTTAAGGTTAGCTTTGATGTTACAGGTTTGGTCATTACACAATTTGGTACATATGACACCAAAAGGGATTAATACTTCCGGGATACACTT													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
CD141_P542	CTGTTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
CD141_P541	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
CD141_P182	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
CD141_P401	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
CD141_P181	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
Consensus	CTATTGATGGATGGAAAGGATTGAGACTTGACTATGGTAACATTACGCGTCCAGACATTCCTTGATAGTGGCAAGGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACGTTGATACGCA													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
CD141_P542	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
CD141_P541	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
CD141_P182	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
CD141_P401	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
CD141_P181	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
Consensus	TGTGAGGAAGGATGGGCCGGAGTTACCCCTATTCTCGTAAATTTGGCTTGATCCTAGTGGAAACAATTTGGTTCATGGCCTGTTCAAGATTAGAACTCTAAGAAAGAAAAGGTCCAATTAAAT													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
CD141_P542	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGATAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
CD141_P541	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGATAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
CD141_P182	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCTCAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGATAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
CD141_P401	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGACAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
CD141_P181	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGATAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
Consensus	AACAAAAAGTTGAACAAAGGAAGAAAGGTTGAATCAAGGAATCACAGTTGCACAGGCTGATGTTGAAGTGATTTTCTATTACACAGTTTGGATAGGCAGAGCCATTTGATCCTAGTTGGGCTGATC													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
CD141_P542	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTCCTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTCCGAATTTTCAAGGC													
CD141_P541	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTCCTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTGAATTTTCAAGGC													
CD141_P182	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTCCTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTGAATTTTCAAGGC													
CD141_P401	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTCCTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTGAATTTTCAAGGC													
CD141_P181	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTACTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTGAATTTTCAAGGC													
Consensus	TTTACGCGCAGAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCCTTTGGGCTACTAATCTTGGCTTCTAAAAACTTAGAAGAATACACACCCGTTTTCTTGAATTTTCAAGGC													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
CD141_P542	TCATGATAAATACAGGTTCTCATGTGTTCCGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
CD141_P541	TCATGATAAATACAGGTTCTTATGTGTTCTGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
CD141_P182	TCATGATAAATACAAAGTTCTTATGTGTTCCGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
CD141_P401	TCATGATAAATACAAAGTTCTTATGTGTTCCGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
CD141_P181	TCATGATAAATACAGGTTCTTATGTGTTCCGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
Consensus	TCATGATAAATACAGGTTCTTATGTGTTCCGATGCCTCAGGTCAGGCCTAAGGAATGAACAACTATGTATAAACCATCATTTGCTGGATATGTAGATGTAGATTTAGCGACAGAGAATTTGTCTCTT													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
CD141_P542	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAATATGCATTACATCAGGGTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACAAATGGCGCGG													
CD141_P541	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAGGGTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGTGTTCAACACAGGCGCGG													
CD141_P182	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAGGGTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACAACTGGCGCGG													
CD141_P401	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAGAGTTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACAACTGGCGCGG													
CD141_P181	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAGGGTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACAACTGGCGCGG													
Consensus	AGAGGTTTGATTGATCATTCGCTAGTGGAAAGTTTTGGTGCTGGAGGAAAAACATGCATTACATCAGGGTTATCCAACTGTTGGCGATATTTGACAAGGCACATTTATTTGCAATTCACAACTGGCGCGG													
	1691	1700	1710	1720	1730	1740	1749							
CD141_P542	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
CD141_P541	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
CD141_P182	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
CD141_P401	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
CD141_P181	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													
Consensus	AGAGAATCACAATTGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAGTTGCACTAG													

[illegible]

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Gene30	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P401	MDCLKKSSLSLPIFLLYFSIILSFNNGVNAASHKVFPGLQSTSTVDYKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYWAHSVSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
Consensus	MDCLKKSSLSLPIFLLYFSIILSFNNGVNAASHKVFPGLQSTSTVDYKNVHRTGYHFQPPKNWINDPNAPHYYNGVYHLFYQYNPYGSVMGNIYWAHSVSTDLINWIPLEPGIYPSEVFDKYGTWSGSAT													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Gene30	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P401	ILPNNKPIILYTGIVDAKNTQVQNYAIPANISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDGYWRTLIGSVHKEQGLAILYKSKNFMKNTKYQHPLHSAADGTGNWECPDFFPYLLHGTNGL													
Consensus	ILPNNKPIILYTGIVDAKNTQVQNYAIPADISDPFLRKWIKPDNNPLIYADVSINKTQFRDPTTCWLGDGYWRTLIGSVHGKQGLAILYKSKNFMKNTKIQHPLHSVDGTGNWECPDFFPYLLHGTNGL													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Gene30	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P401	DASYNKKNIKHVLYKSLDVTRFEYTYVGKYDTKKDRIYIPDKTSIDGWKGLRLDYGNYYASKSFYDPSKNRRIMHGWANESDTYNDVYKKGWAGIQTSPRKLWLDPSGKQLYQWPVEELETIRENKIQLM													
Consensus	DASYNKKNIKHVLYKSLDVTRFEYTYVGIYDTKKDRIYIPDKTSIDGWKGLRLDYGNYYASKSFYDPSKNRRIMHGWANESDTYNDVYKKGWAGIQTIPRKLWLDPSGKQLYQWPVEELETIRENKIQLM													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Gene30	-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P401	RKLNGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPNMANLYAQDYCAIKGSTVQGGGLGPFGLLTLASQNLEETPYVFRVFKAQDKYKYLHCSDATRSTLKNOKTHYKPSFAGYVDVDLTNKTLSLR													
Consensus	RKLNGDKIEVKGITPAQADVEYTFSSSLDKAEPFDPNMANLYAQDYCAIKGSTVQGGGLGPFGLLTLASQNLEETPYVFRVFKAQDKYKYLHCSDASRSTLKNOKTHYKPSFAGYVDVDLTNKTLSLR													
	521	530	540	550	560	570	580	590						
Gene30	-----+-----+-----+-----+-----+-----+-----+-----													
CD111_P401	SLIDHSVYESFGAGGKTCITSRVYPTLAIYDNAHLFVFNNGTETIKIKSLNAWTHGPKKNWSFGHSSFR													
Consensus	SLIDHSVYESFGAGGKTCITSRVYPTLAIYDNAHLFVFNNGTETIKIKSLNAWTHGPKKNWSFGHSSFR													

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Gene30	ATGATTGTTAAAAAAGCTCTCTCTTTTTCTTTGGCAATTTTTTGTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGGTGCATCTACTAGCA													
CD111_P401	ATGGATTGTTAAAAAAGCTCTCTCTTTTTCTTTTCTTTGCCAATTTTTTGTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGGTGCATCTACAGCA													
Consensus	ATGGATTGTTAAAAAAGCTCTCTCTTTTTCTTTGCCAaTTTTTTGTATATTTCTCCATAATTTTATCTTTCAACAATGGCGTTAATGCTTCACACAAGTTTTTCAGGGGTGCATCTACaAGCA													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Gene30	CGGTTGATGTGA AAAATGTTTCATAGAACTGGGTATCATTTTTCACACCTCCTAAAAACTGGGATTATGGTACGTTCACGTTCTTTCCACACCTTCATCTATATAAATATTGTATGATTTAAGTTATACATGTTG													
CD111_P401	CGGTTGATGTGA AAAATGTTTCATAGAACTGGGTATCATTTTTCACACCTCCTAAAAACTGGGATCATGCTGATGTTTTCACACCTTCATCTATATAAATATTGTATGATTTAAGTTATACATGTTG													
Consensus	CGGTTGATGTGA AAAATGTTTCATAGAACTGGGTATCATTTTTCACACCTCCTAAAAACTGGGATcATG.....													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Gene30	ATAGTATATTTTTTTTTATGTTATTAGTACATAGTATATCTTAAATTACATGTAAAGACACGTTAATTAGTCACTGTATCTCATAAATGACCTAGTTATGATTTTGTGGTGCATATATGTATGTGAG													
CD111_P401	ATAGTATATTTTTTTTTATGTTATTAGTACATAGTATATCTTAAATTACATGTAAAGACACGTTAATTAGTCACTGTATCTCATAAATGACCTAGTTATGATTTTGTGGTGCATATATGTATGTGAG													
Consensus													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Gene30	ATAGATAGATCATTTGCATTCATGATTATGACCTCATGATTTTTATCTTTTTGCTATGTTTCTGAATAACTAGATTTTGTTTATTCAGTTAATTTGGACGTTGACTGGAAATATTATTATTCAGTTA													
CD111_P401	ATAGATAGATCATTTGCATTCATGATTATGACCTCATGATTTTTATCTTTTTGCTATGTTTCTGAATAACTAGATTTTGTTTATTCAGTTAATTTGGACGTTGACTGGAAATATTATTATTCAGTTA													
Consensus													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Gene30	TTTTAATTTCTTACTCTGATGCGTTTTTGATTTAATGTCTATCGAAAACAGTCTTTCTACCTCCTCGAGGTAATGATAGGCTCTATGTACGTTTAACTCTCCGAATTCGTTTTAGTAGATATGTTATTG													
CD111_P401	TTTTAATTTCTTACTCTGATGCGTTTTTGATTTAATGTCTATCGAAAACAGTCTTTCTACCTCCTCGAGGTAATGATAGGCTCTATGTACGTTTAACTCTCCGAATTCGTTTTAGTAGATATGTTATTG													
Consensus													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Gene30	TTATTTTACGGCTTTCGAGTTAATTAACTAGTATAGTAAAAA AAAAAAACAATAAGTATGTGGTTGTGGATGGTTGAGATTTTATCCAGCGCTTAATTAGTGATTTCAAGTTCGACTTGTTAAGGAT													
CD111_P401	TTATTTTACGGCTTTCGAGTTAATTAACTAGTATAGTAAAAA AAAAAAACAATAAGTATGTGGTTGTGGATGGTTGAGATTTTATCCAGCGCTTAATTAGTGATTTCAAGTTCGACTTGTTAAGGAT													
Consensus													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Gene30	GAAATAAATATATGTTGGAGCGCTCAATTTTAAAGATGGTCACTGCAATGTATGAATCAAAATCAATCGTGTCTCAACGTAATATCGAACATAAAGGAGGAAAAAGATGAACACTACGTGATATCCA													
CD111_P401	GAAATAAATATATGTTGGAGCGCTCAATTTTAAAGATGGTCACTGCAATGTATGAATCAAAATCAATCGTGTCTCAACGTAATATCGAACATAAAGGAGGAAAAAGATGAACACTACGTGATATCCA													
Consensus													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Gene30	TACCTTTTTTGGTAGTACTTGTGGTGTACTGCAAGGATATCTTAAACCTATGTCTACTAAGTTATCGCTGGGGACCCATATCTCTGGGGGCCATCTAAACAACAACAACACTACTTATATATGTTATT													
CD111_P401	TACCTTTTTTGGTAGTACTTGTGGTGTACTGCAAGGATATCTTAAACCTATGTCTACTAAGTTATCGCTGGGGACCCATATCTCTGGGGGCCATCTAAACAACAACAACACTACTTATATATGTTATT													
Consensus													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Gene30	TAGAAAAA A A A A A AATATATGTA CTATGAGTGTATGTTCTACTAAGGA A AATGCTTATGAGATGAATATAACCTGAATTTCTGAAGAACA AATTTTGAGAAATATGAATAACGAATCACAAAGTC													
CD111_P401	TAGAAAAA A A A A A AATATATGTA CTATGAGTGTATGTTCTACTAAGGA A AATGCTTATGAGATGAATATAACCTGAATTTCTGAAGAACA AATTTTGAGAAATATGAATAACGAATCACAAAGTC													
Consensus													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Gene30	CCTTAAAGATCTTTTAGAGTTGATTACATTATATATGTAGAGTAGATCAAACTATTTTTTCACTTCTAATAAGACTTGAACCAATTTGAGCATATGA A AATATTTTTTGATTCTTATTTAATTGCAAA													
CD111_P401	CCTTAAAGATCTTTTAGAGTTGATTACATTATATATGTAGAGTAGATCAAACTATTTTTTCACTTCTAATAAGACTTGAACCAATTTGAGCATATGA A AATATTTTTTGATTCTTATTTAATTGCAAA													
Consensus													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Gene30	ATATTGAGATCACTAGATGATGAGAATTTAAGAAATATTTGCTTAACCTTGAACCTTTCTTAAACATGCATAATAGTTATTCTGTTTTGATGGTTTAAATTTATTTTTGAAATAAAGTATTGTTAA													
CD111_P401	ATATTGAGATCACTAGATGATGAGAATTTAAGAAATATTTGCTTAACCTTGAACCTTTCTTAAACATGCATAATAGTTATTCTGTTTTGATGGTTTAAATTTATTTTTGAAATAAAGTATTGTTAA													
Consensus													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Gene30	GAGAAATAATACAGTAGAGTAATGATCTAATGA A AATACTCAATCAATAA A AAGACTTGATTCCTTTTCCAATGCACATACTGCTTATAGAATAATGTTAACAGTTCATGTAACCGTTGCTCAACCC													
CD111_P401	GAGAAATAATACAGTAGAGTAATGATCTAATGA A AATACTCAATCAATAA A AAGACTTGATTCCTTTTCCAATGCACATACTGCTTATAGAATAATGTTAACAGTTCATGTAACCGTTGCTCAACCC													
Consensus													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690

	1	10	20	30	40	50	60	70	80	90	100	110	120	130
Gene40	ATGGAGATTTTAGAAATATCTTCTCTCTTTGGGCTTTGCCAATTCCTTTTGTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACCTTGCACCTTACTACTAGCC													
CD141_P401	ATGGAGATTTTAGAAAGATCTTCTCTCTCTTTGGGCTTTGCCAATTCCTTTTGTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACCTTGCACCTTACTACTAGCC													
Consensus	ATGGAGATTTTAGAAATATCTTCTCTCTTTGGGCTTTGCCAATTCCTTTTGTGTGCTTCTTATCAACAATGGAGTATTTGTTGATGCTTCTCACAAAGTTTATATGCACCTTGCACCTTACTACTAGCC													
	131	140	150	160	170	180	190	200	210	220	230	240	250	260
Gene40	ATGTTGATGCTAGCAAGGTCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATTACGGTATGTTTTTCATTGTTTTTCATGCTATGTTTCTGTCTCAATTTATGATTAGACGTTCTGA													
CD141_P401	ATGTTGATGCTAGCAAGGTCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATTACGGTATGTTTTTCATTGTTTTTCATGCTATGTTTCTGTCTCAATTTATGATTAGACGTTCTGA													
Consensus	ATGTTGATGCTAGCAAGGTCATAGAACTGGTTATCATTTTCAACCTCCTAAAACTGGATTACGG.....													
	261	270	280	290	300	310	320	330	340	350	360	370	380	390
Gene40	GTTCTAAGTTGAACATAAAAGTTTTGAAATATGTATTTATCTTATTGTTGTTTGGTTTTTGTACAAATTATATGAAATGATGTTTGGGTTAAAGAGTCATGATTTTGCCAAATTTATTGTTAGGGGC													
CD141_P401	GTTCTAAGTTGAACATAAAAGTTTTGAAATATGTATTTATCTTATTGTTGTTTGGTTTTTGTACAAATTATATGAAATGATGTTTGGGTTAAAGAGTCATGATTTTGCCAAATTTATTGTTAGGGGC													
Consensus													
	391	400	410	420	430	440	450	460	470	480	490	500	510	520
Gene40	GGAGCTAAAGGTTAAGTACGGCCATAGAGGTTCTGCCAATGCAACAGTATTAGTTTGATTGTTGTTATTTATATTAGAAATTAACCTAAGTAGGTATACAAAGTAATTTAGACTCAATTACTATATAC													
CD141_P401	GGAGCTAAAGGTTAAGTACGGCCATAGAGGTTCTGCCAATGCAACAGTATTAGTTTGATTGTTGTTATTTATATTAGAAATTAACCTAAGTAGGTATACAAAGTAATTTAGACTCAATTACTATATAC													
Consensus													
	521	530	540	550	560	570	580	590	600	610	620	630	640	650
Gene40	AATATGATCAGTATTTTTAAATTTAGARTAACTTAAAGGTTGCATCCGCTTTTGTCCGTTGCTTTTCTGTGCTTCGGGAAGTATCTACATATGTATTAATTACTTATTTGTGGATAAATAAAATGAAC													
CD141_P401	AATATGATCAGTATTTTTAAATTTAGARTAACTTAAAGGTTGCATCCGCTTTTGTCCGTTGCTTTTCTGTGCTTCGGGAAGTATCTACATATGTATTAATTACTTATTTGTGGATAAATAAAATGAAC													
Consensus													
	651	660	670	680	690	700	710	720	730	740	750	760	770	780
Gene40	AGAGCAAAATTTAGAAATGATTTATTTTAATATTATTTCCAGCTGATTGGTTAGGTTGACTGAAATGTTAGGCTATTTTGATACCACATGTTATAAATAATATAATAATAATAATACTCTTGACAG													
CD141_P401	AGAGCAAAATTTAGAAATGATTTATTTTAATATTATTTCCAGCTGATTGGTTAGGTTGACTGAAATGTTAGGCTATTTTGATACCACATGTTATAAATAATATAATAATAATAATACTCTTGACAG													
Consensus													
	781	790	800	810	820	830	840	850	860	870	880	890	900	910
Gene40	TTTTTGAGTTAATGAACCTACATGTTGCCCATGATGAGCAGATTACATATGAATAATTAGTATCGTAATTAAATATTGCCGTTCTATAGATATTATTAAATAATATATACAAACGAGTAAGATTGAA													
CD141_P401	TTTTTGAGTTAATGAACCTACATGTTGCCCATGATGAGCAGATTACATATGAATAATTAGTATCGTAATTAAATATTGCCGTTCTATAGATATTATTAAATAATATATACAAACGAGTAAGATTGAA													
Consensus													
	911	920	930	940	950	960	970	980	990	1000	1010	1020	1030	1040
Gene40	AATATTTTTTGTGTTGTTACATCATCAACTTATAGTCAAACTTAGGTGCCAGCCTGCTATTTTTTACTTATACAATAATACTAATTTTTTTATTGAAATATTAACTCTAATATGATAGGAAATGAT													
CD141_P401	AATATTTTTTGTGTTGTTACATCATCAACTTATAGTCAAACTTAGGTGCCAGCCTGCTATTTTTTACTTATACAATAATACTAATTTTTTTATTGAAATATTAACTCTAATATGATAGGAAATGAT													
Consensus													
	1041	1050	1060	1070	1080	1090	1100	1110	1120	1130	1140	1150	1160	1170
Gene40	CGTATAAGTTAGAAAAAGAAATATTGAGGTACAACTAGTAATATTCTTTCATTCTTAATCAGATGTCTCGAGTTTGAGTCTACTTGAGTACGAAGTCATTTTTGTTAGGAAGCATTTCACCCAATGT													
CD141_P401	CGTATAAGTTAGAAAAAGAAATATTGAGGTACAACTAGTAATATTCTTTCATTCTTAATCAGATGTCTCGAGTTTGAGTCTACTTGAGTACGAAGTCATTTTTGTTAGGAAGCATTTCACCCAATGT													
Consensus													
	1171	1180	1190	1200	1210	1220	1230	1240	1250	1260	1270	1280	1290	1300
Gene40	GGAACTTTTGGGCGTGAATCTCGATTTAGTCGGGCTCACCAGACCCGGATGAAAAAAGAAATTAATATAGTATATATAGTATAGTCAACTCCACCTCCCATCCCCAAACCGAGTAGGGTACGTTGG													
CD141_P401	GGAACTTTTGGGCGTGAATCTCGATTTAGTCGGGCTCACCAGACCCGGATGAAAAAAGAAATTAATATAGTATATATAGTATAGTCAACTCCACCTCCCATCCCCAAACCGAGTAGGGTACGTTGG													
Consensus													
	1301	1310	1320	1330	1340	1350	1360	1370	1380	1390	1400	1410	1420	1430
Gene40	GTTGGTGGTAGAATCTGAACCAATCGTATATATACATTATTTATATTATAGTATACAAATAAAACATGTGAAGAGCCCAAAGTACGTGAATATTATAGAGCCAAATATATGACGTAAAGATGAAGAAG													
CD141_P401	GTTGGTGGTAGAATCTGAACCAATCGTATATATACATTATTTATATTATAGTATACAAATAAAACATGTGAAGAGCCCAAAGTACGTGAATATTATAGAGCCAAATATATGACGTAAAGATGAAGAAG													
Consensus													
	1431	1440	1450	1460	1470	1480	1490	1500	1510	1520	1530	1540	1550	1560
Gene40	AAATTTAAATAAAATATGAATTACTCTATACATTCGTTATGATTTTCATTTGTTCTCTAGACAATTCAGTTTGATATATTTTAAGAAAAAAGGATAGACATAAATAAGTAAATAATATATATACAAATA													
CD141_P401	AAATTTAAATAAAATATGAATTACTCTATACATTCGTTATGATTTTCATTTGTTCTCTAGACAATTCAGTTTGATATATTTTAAGAAAAAAGGATAGACATAAATAAGTAAATAATATATATACAAATA													
Consensus													
	1561	1570	1580	1590	1600	1610	1620	1630	1640	1650	1660	1670	1680	1690
Gene40	ATAAATATAGTCCCTTGTGTTACTATCATGATAAATAGAAAAAATAAATTAGAAACGACCTTTTTTTTTTTTGGGGGGGGGAGGGGGGAGCGGGGGTTATATTGAGGAATAAAGAAAGTCAAAA													
CD141_P401	ATAAATATAGTCCCTTGTGTTACTATCATGATAAATAGAAAAAATAAATTAGAAACGACCTTTTTTTTTTTGGGGGGGGGAGGGGGGAGCGGGGGTTATATTGAGGAATAAAGAAAGTCAAAA													
Consensus													
	1691	1700	1710	1720	1730	1740	1750	1760	1770	1780	1790	1800	1810	1820
Gene40	TGTTTTTGACAAATATTGGCAACTTGTAGCCTGTAGGAAGAGCTTTGCTTCGGGCACATCCTTCTGTTGCTCCGACACAGCGGATTCAGGATTTTAAATTTATGAGACTCCATAGTAGAAATGATT													
CD141_P401	TGTTTTTGACAAATATTGGCAACTTGTAGCCTGTAGGAAGAGCTTTGCTTCGGGCACATCCTTCTGTTGCTCCGACACAGCGGATTCAGGATTTTAAATTTATGAGACTCCATAGTAGAAATGATT													
Consensus													
	1821	1830	1840	1850	1860	1870	1880	1890	1900	1910	1920	1930	1940	1950
Gene40	CTAAAAATTTGAGTATGAGTTCATGTAACCTATAACATTTATTCAGACTCTATATATAGTTTTTAAAAATCTATTAATATTTGATAGTGAACACGCTCTTGTTATTATTATAGTATTTATACCTTGTAGTC													
CD141_P401	CTAAAAATTTGAGTATGAGTTCATGTAACCTATAACATTTATTCAGACTCTATATATAGTTTTTAAAAATCTATTAATATTTGATAGTGAACACGCTCTTGTTATTATTATAGTATTTATACCTTGTAGTC													
Consensus													
	1951	1960	1970	1980	1990	2000	2010	2020	2030	2040	2050	2060	2070	2080
Gene40	ATCGTAGGAACTCATCTAGATCTATTAGTATCTATAAATTTTCGCTTATTAATTTCTGAATGAATATTATTATACATATTAAATATTTTCAGATAAGCTATTGAGTTTGTGTTACTCGTAACTTCAA													
CD141_P401	ATCGTAGGAACTCATCTAGATCTATTAGTATCTATAAATTTTCGCTTATTAATTTCTGAATGAATATTATTATACATATTAAATATTTTCAGATAAGCTATTGAGTTTGTGTTACTCGTAACTTCAA													
Consensus													
	2081	2090	2100	2110	2120	2130	2140	2150	2160	2170	2180	2190	2200	2210
Gene40	TAGTATATTGAGTTTTCTTTTGTATAGTCATAAATTTCTAGTTTCTCTTTAATTTGCCCTTTTTCCATGGAAAGAGGTTGTGGGCTTGCATGAGTTGGTTCCCATTAGAACTTACAAAAAAA													
CD141_P401	TAGTATATTGAGTTTTCTTTTGTATAGTCATAAATTTCTAGTTTCTCTTTAATTTGCCCTTTTTCCATGGAAAGAGGTTGTGGGCTTGCATGAGTTGGTTCCCATTAGAACTTACAAAAAAA													
Consensus													
	2211	2220	2230	2240	2250	2260	2270	2280	2290	2300	2310	2320	2330	2340
Gene40	AAAAAAAAAAAAAGGTAGATTCTATGTTGAAAAAGAAAAAAAACGCCTAARACTTTTTATATTTCCTAGATTTTTTGTATTTTAGTGACAGTGACAGCACATTCATCAAACTTCAAAATATGTGC													
CD141_P401	AAAAAAAAAAAAAGGTAGATTCTATGTTGAAAAAGAAAAAAAACGCCTAARACTTTTTATATTTCCTAGATTTTTTGTATTTTAGTGACAGTGACAGCACATTCATCAAACTTCAAAATATGTGC													
Consensus													
	2341	2350	2360	2370	2380	2390	2400	2410	2420	2430	2440	2450	2460	2470
Gene40	ATTAATTTATCATCTAATATTTATTAGCTGGATGATCCATGTACTATTCTTCATCCCTTAGGTTTGATTTTACATTTTACAAATTATAGTTTAAATTTCTTGCTAACATTTAATTATGTTTGGAC													
CD141_P401	ATTAATTTATCATCTAATATTTATTAGCTGGATGATCCATGTACTATTCTTCATCCCTTAGGTTTGATTTTACATTTTACAAATTATAGTTTAAATTTCTTGCTAACATTTAATTATGTTTGGAC													
Consensus													
	2471	2480	2490	2500	2510	2520	2530	2540	2550	2560	2570	2580	2590	2600
Gene40	TAATCAAACTCARTTGTTCCTTTTTTTCATGTTTCTTCATGATGCTTCAACAACAACAGATCCCAATGGTGAAGTCAAAATTTATTTTTCACAAATCTTTTACTAGCCATAATTAATTTTA													
CD141_P401	TAATCAAACTCARTTGTTCCTTTTTTTCATGTTTCTTCATGATGCTTCAACAACAACAGATCCCAATGGTGAAGTCAAAATTTATTTTTCACAAATCTTTTACTAGCCATAATTAATTTTA													
ConsensusATCCAAATGG.....ATCCAAATGG.....C.....													
	2601	2610	2620	2630	2640	2650	2660	2670	2680	2690	2700	2710	2720	2730
Gene40	TTTACTTTGTAGTAATTTTTTTGTGCTTTTTGTTCTCAATAGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGGAGCGATATGGGGCAACATTATTGGGCTCAT													
CD141_P401	TTTACTTTGTAGTAATTTTTTTGTGCTTTTTGTTCTCAATAGGTCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGGAGCGATATGGGGCAACATTATTGGGCTCAT													
ConsensusCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGGAGCGATATGGGGCAACATTATTGGGCTCATCCAATGTATTACATGGAGTGTATCATTTATTCTACCAAGTACACCCAAAGGGAGCGATATGGGGCAACATTATTGGGCTCAT													
	2731	2740	2750	2760	2770	2780	2790	2800	2810	2820	2830	2840	2850	2860
Gene40	CGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATTACCCGTCCAAGATATTTGACAGATATGGTACATGGTCCGGGTCAGCCCAATCTTGCCAGGCCAACAGGCTGTTATTCTCTA													
CD141_P401	CGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATTACCCGTCCAAGATATTTGACAGATATGGTACATGGTCCGGGTCAGCCCAATCTTGCCAGGCCAACAGGCTGTTATTCTCTA													
ConsensusCGGTCTCAAGGACTTGATCAATTGGATCCCACTTGAACCCGCTATTACCCGTCCAAGATATTTGACAGATATGGTACATGGTCCGGGTCAGCCCAATCTTGCCAGGCCAACAGGCTGTTATTCTCTA													
	2861	2870	2880	2890	2900	2910	2920	2930	2940	2950	2960	2970	2980	2990
Gene40	CACCGGAATTTGGATGCTAATAAGACACAGTCCAAATTTATGCARTCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAGGCCGATACCAATCCATTGATTGTTGCTGATAAAACCATC													
CD141_P401	CACCGGAATTTGGATGCTAATAAGACACAGTCCAAATTTATGCARTCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAGGCCGATACCAATCCATTGATTGTTGCTGATAAAACCATC													
ConsensusCACCGGAATTTGGATGCTAATAAGACACAGTCCAAATTTATGCARTCCGGCTAACATGTCTGATCCATATCTTCGTAAGTGGATCAGGCCGATACCAATCCATTGATTGTTGCTGATAAAACCATC													
	2991	3000	3010	3020	3030	3040	3050	3060	3070	3080	3090	3100	3110	3120
Gene40	AACAAAGCCCAATTTCTGATCCACACACAGCATGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTGGGAGTGTAGGAATCATAGGGGAAGGTTAATGTACAAAGTATATAGAACTTCATGA													
CD141_P401	AACAAAGCCCAATTTCTGATCCACACACCGCATGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTAGGAGTGTAGGAATCATAGGGGAAGGTTAATGTACAAAGTATATAGAACTTCATGA													
ConsensusAACAAAGCCCAATTTCTGATCCACACACAGCATGGATGGGCCGAGATGGAATTTGGAGATCTTGGTTAGGAGTGTAGGAATCATAGGGGAAGGTTAATGTACAAAGTATATAGAACTTCATGA													
	3121	3130	3140	3150	3160	3170	3180	3190	3200	3210	3220	3230	3240	3250
Gene40	AATGGGACCAAGCCCAACACCCACTCCACTCAGCTCCCGGTACTTGGAAATTTGGGATGTCTCGATTTTTCCTAGTGTCTAATAAATAAAGATGGTTTGGACAGCTCATCAATGGCAAGACATTAA													
CD141_P401	AATGGGACCAAGCCCAACACCCACTCCACTCAGCTCCCGGTACTTGGAAATTTGGGATGTCTCGATTTTTCCTAGTGTCTAATAAATAAAGATGGTTTGGACAGCTCATCAATGGCAAGACATTAA													
ConsensusAATGGGACCAAGCCCAACACCCACTCCACTCAGCTCCCGGTACTTGGAAATTTGGGATGTCTCGATTTTTCCTAGTGTCTAATAAATAAAGATGGTTTGGACAGCTCATCAATGGCAAGACATTAA													
	3251	3260	3270	3280	3290	3300	3310	3320	3330	3340	3350	3360	3370	3380
Gene40	ACATGTTCTTAAGATTAGCTTTGATGTTACTAGGTTTGATCATTTACACAAATTTGGTACATATGACACCAAAAGGATAGTACTTTCCGGATAACACTTCTATTGATGGATGGAAGGATTGAGACTTGAC													
CD141_P401	ACATGTTCTTAAGATTAGCTTTGATGTTACTAGGTTTGATCATTTACACAAATTTGGTACATATGACACCAAAAGGATAGTACTTTCCGGATAACACTTCTATTGATGGATGGAAGGATTGAGACTTGAC													
ConsensusACATGTTCTTAAGATTAGCTTTGATGTTACTAGGTTTGATCATTTACACAAATTTGGTACATATGACACCAAAAGGATAGTACTTTCCGGATAACACTTCTATTGATGGATGGAAGGATTGAGACTTGAC													
	3381	3390	3400	3410	3420	3430	3440	3450	3460	3470	3480	3490	3500	3510
Gene40	TATGGTAACATTACCGCTCCAGACATCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACCTCGTGATACGATGTGAGGAAGAGGTTGGGCGGGAGTTACCCCTA													
CD141_P401	TATGGTAACATTACCGCTCCAGACATCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACCTCGTGATACGATGTGAGGAAGAGGTTGGGCGGGAGTTACCCCTA													
ConsensusTATGGTAACATTACCGCTCCAGACATCTTTGATAGTGGCAAGATCGTAGGATTTTGTGGGTTGGGCTAATGAATCAGATACCTCGTGATACGATGTGAGGAAGAGGTTGGGCGGGAGTTACCCCTA													
	3511	3520	3530	3540	3550	3560	3570	3580	3590	3600	3610	3620	3630	3640
Gene40	TTCTCGTAAATTTGGCTTGATCTAGTGGAACAATTTGGTTCAATGGCTGTTTCAGAAATTAGAACTCTAGAAAGAAAAAGGTTCAATTAATAACAAAAGTTGAACAGGGAGAAAGGTTGA													
CD141_P401	TTCTCGTAAATTTGGCTTGATCTAGTGGAACAATTTGGTTCAATGGCTGTTTCAGAAATTAGAACTCTAGAAAGAAAAAGGTTCAATTAATAACAAAAGTTGAACAGGGAGAAAGGTTGA													
ConsensusTTCTCGTAAATTTGGCTTGATCTAGTGGAACAATTTGGTTCAATGGCTGTTTCAGAAATTAGAACTCTAGAAAGAAAAAGGTTCAATTAATAACAAAAGTTGAACAGGGAGAAAGGTTGA													
	3641	3650	3660	3670	3680	3690	3700	3710	3720	3730	3740	3750	3760	3770
Gene40	AATCAAGGAATCAGAGTTGCACAGGTTTGGGCTTGATTCTCTCTGTTTTCTTTTGTGTTATACTTATTGAGACGCATGTGAARAAGCAATTAATTTGAATTTGCAGGCTGATGTTGAGTGATT													
CD141_P401	AATCAAGGAATCAGAGTTGCACAGGTTTGGGCTTGATTCTCTCTGTTTTCTTTTGTGTTATACTTATTGAGACGCATGTGAARAAGCAATTAATTTGAATTTGCAGGCTGATGTTGAGTGATT													
ConsensusAATCAAGGAATCAGAGTTGCACAGG.....CTGATGTTGAGTGATT.....CTGATGTTGAGTGATT													
	3771	3780	3790	3800	3810	3820	3830	3840	3850	3860	3870	3880	3890	3900
Gene40	TTCTCATTCGAAAGTTTGGAAAGGACAGCTATTTTGATCCTAGTTGGGCTGATCTTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCTTTTGGGCTCTTACTT													
CD141_P401	TTCTCATTCGAAAGTTTGGAAAGGACAGCTATTTTGATCCTAGTTGGGCTGATCTTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCTTTTGGGCTCTTACTT													
ConsensusTTCTCATTCGAAAGTTTGGAAAGGACAGCTATTTTGATCCTAGTTGGGCTGATCTTTACGCGCAAGATGTATGCGCCATTAAAGGTTCAACGGTCCAGGTTGGACTTGGGCTTTTGGGCTCTTACTT													
	3901	3910	3920	3930	3940	3950	3960	3970	3980	3990	4000	4010	4020	4030
Gene40	TGCTCTCAAAAACCTTAGAAGATACACACCCGTTTTCTTGAATTTCAAGGCTCAGATAAATACAAAGTTCTTATGTGTTCCGATGCCCTCAGGTTTGTTGTTCTTCATTTTTAACGCCCTCGCCCA													
CD141_P401	TGCTCTCAAAAACCTTAGAAGATACACACCCGTTTTCTTGAATTTCAAGGCTCAGATAAATACAAAGTTCTTATGTGTTCCGATGCCCTCAGGTTTGTTGTTCTTCATTTTTAACGCCCTCGCCCA													
ConsensusTGCTCTCAAAAACCTTAGAAGATACACACCCGTTTTCTTGAATTTCAAGGCTCAGATAAATACAAAGTTCTTATGTGTTCCGATGCCCTCAGGTTTGTTGTTCTTCATTTTTAACGCCCTCGCCCA													
	4031	4040	4050	4060	4070	4080	4090	4100	4110	4120	4130	4140	4150	4160
Gene40	CCCCAAAAAAGAAAGACATGTGCATTTGACAAATATTGTTATTCAACTTCAATTGAATGTTTGTGTTATTGTTTCCATTGATGTCCACAGGTCAGGCCTAAGAGATGAACCACTATGTATAAACCATCA													
CD141_P401	CCCCAAAAAAGAAAGACATGTGCATTTGACAAATATTGTTATTCAACTTCAATTGAATGTTTGTGTTATTGTTTCCATTGATGTCCACAGGTCAGGCCTAAGAGATGAACCACTATGTATAAACCATCA													
ConsensusCCCCAAAAAAGAAAGACATGTGCATTTGACAAATATTGTTATTCAACTTCAATTGAATGTTTGTGTTATTGTTTCCATTGATGTCCACAGGTCAGGCCTAAGAGATGAACCACTATGTATAAACCATCA													
	4161	4170	4180	4190	4200	4210	4220	4230	4240	4250	4260	4270	4280	4290
Gene40	TTTGCTGGATATGTAGATGTAGATTTAGCAACACAGAATTTGCTCTTGAAGTTTGTAGGCCCTCATCTCAATTCCTTTGTAATTTTACTATTTCATATAAATTTTCGCGTGTTTTAACAAATTTGGTTA													
CD141_P401	TTTGCTGGATATGTAGATGTAGATTTAGCAACACAGAATTTGCTCTTGAAGTTTGTAGGCCCTCATCTCAATTCCTTTGTAATTTTACTATTTCATATAAATTTTCGCGTGTTTTAACAAATTTGGTTA													
ConsensusTTTGCTGGATATGTAGATGTAGATTTAGCAACACAGAATTTGCTCTTGAAGTTTGTAGGCCCTCATCTCAATTCCTTTGTAATTTTACTATTTCATATAAATTTTCGCGTGTTTTAACAAATTTGGTTA													
	4291	4300	4310	4320	4330	4340	4350	4360	4370	4380	4390	4400	4410	4420
Gene40	TATTTCCAGATTGATCATTTCGGTAGTGGAAAGTTTGGTGCTGGAGGAAACATGTCATTACATCARGAGTTTATCCAACTTTGGCGATATTTGACAAAGCACTTTATTTGGCTTCAACAAATGGGCGGA													
CD141_P401	TATTTCCAGATTGATCATTTCGGTAGTGGAAAGTTTGGTGCTGGAGGAAACATGTCATTACATCARGAGTTTATCCAACTTTGGCGATATTTGACAAAGCACTTTATTTGGCTTCAACAAATGGGCGGA													
ConsensusATTGATCATTTCGATAGTGGAAAGTTTGGTGCTGGAGGAAACATGTCATTACATCARGAGTTTATCCAACTTTGGCGATATTTGACAAAGCACTTTATTTGGCTTCAACAAATGGGCGGA													
	4421	4430	4440	4450	4460	4470	4478							
Gene40	GAGAACTCAAAATGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTGCACATAG													
CD141_P401	GAGAACTCAAAATGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTGCACATAG													
ConsensusGAGAACTCAAAATGAGACTCTAAATGCTTGGAGCATGGCTAATGCAAAAGTGCACATAG													